UAC

SEARCH REQUEST FORM

Scientific and Technical Information Center

Art Unit: 1623 Phone Mail Box and Bldg/Room Location CM 8B19 CM 8A0 If more than one search is subm	itted, please prioritiz	*********	PAPER DISK E-MAIL d. *********************************
Please provide a detailed statement of the Include the elected species or structures, k utility of the invention. Define any terms known. Please attach a copy of the cover statement of the statement of the statement of the Include Statement of th	eywords, synonyms, acron that may have a special me	yms, and registry numbers, and cor aning. Give examples or relevant	nbine with the concept or
Title of Invention:		- A	
Inventors (please provide full names): _	see atta	asse home	heet
Earliest Priority Filing Date:	9-25-0	/	
glucosa per cl uomig	search a mine an 1-13, a it to al	·	comprising algesic dof ain per.
Jan Delaval Reference Librar Biolechnology & Chemic CA1 1E07 – 703-308 jan delaval@uspto	ian eal Library i-4498	hanks.	*******
STAFF USE ONLY	Type of Search	Vendors and cost where	e applicable
Searcher:	NA Sequence (#)	STN	
Searcher Phone #:	AA Sequence (#)	Dialog	
Searcher Location:	Structure (#)	Questel/Orbit	
Date Searcher Picked Up:(/2/7//3	Bibliographic	Dr.Link	
Date Completed: 1/2 Y/d25	Litigation	Lexis/Nexis	
Searcher Prep & Review Time:	Fulltext	Sequence Systems	
Clerical Prep Time:	. Patent Family	WWW/Internet	

STST AVAILABLE COPY

4)

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(FILE 'HOME' ENTERED AT 10:34:45 ON 27 JAN 2003) SET COST OFF
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FILE 'REGISTRY' ENTERED AT 10:34:56 ON 27 JAN 2003
L1
               3 S 3416-24-8 OR 29031-19-4 OR 66-84-2
                 E .ALPHA.-GLUCOSAMINE/CN
L2
               1 S E4
L3
               1 S 6490-70-6
                 E .BETA.-GLUCOSAMINE/CN
T.4
               1 S E4
L5
               1 S 14257-69-3
                 E N-ACETYL-D-GLUCOSAMINE/CN
L6
               1 S E3
T.7
             297 S (7512-17-6 OR 3416-24-8 OR 6490-70-6 OR 14257-69-3)/CRN
\Gamma8
              37 S L7 AND (7664-93-9/CRN OR CLH)
L9
              8 S L8 AND 2/NC
L10
              12 S L1-L6, L9
L11
              1 S IBUPROFEN/CN
L12
              1 S KETOPROFEN/CN
L13
              18 S C13H18O2/MF AND 46.150.18/RID AND 1/NR AND BENZENEACETIC AND
L14
              13 S L13 AND 2 METHYLPROPYL
L15
              3 S L14 AND IBUPROFEN
L16
              15 S L13 NOT L15
L17
              12 S C16H14O3/MF AND 46.150.18/RID AND 2/NR AND BENZENEACETIC AND
L18
               3 S L17 AND KETOPROFEN
L19
               9 S L17 NOT L18
L20
               6 S L11, L12, L15, L18
                 SEL RN
L21
             426 S E1-E6/CRN
L22
               2 S L21 AND L7
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L23
            9874 S L10
           27519 S ?GLUCOSAMINE? OR ACETYLGLUCOSAMINE OR ACETYL (1W) GLUCOSAMINE
L24
L25
          29352 S L23, L24
L26
            8313 S L20
L27
           8906 S IBUPROFEN OR KETOPROFEN
L28
          10023 S L26, L27
L29
           4491 S NSAID
L30
          11691 S (NONSTEROID? OR NON STEROID?) (L) ?INFLAM?
          49473 S ANALGES?
L31
                 E ANALGESIC/CT
                 E E6+ALL
L32
          27328 S E5
L33
          54026 S E5+NT
                 E E22+ALL
L34
          17760 S E5+NT
                 E ANTIINFLAM/CT
                 E E5+ALL
                                                            Jan Delaval
L35
          19798 S E2
                                                         Reference Librarian
                 E E2+ALL
                                                    Biotechnology & Chemical Library
          24177 S E4,E5
                                                       CM1 1E07 - 703-308-4498
L37
          28056 S E3+NT
                                                         jan.delaval@uspto.gov
L38
             36 S L25 AND L28
L39
            329 S L25 AND L29-L37
L40
             23 S L38 AND L39
L41
              36 S L38, L40
                 E ANTIARTHRITIC/CT
                 E E4+ALL
L42
           4488 S E5, E4+NT
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E ANTIHISTAMIN/CT

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L43
            1153 S E5-E7
                 E E4+ALL
L44
            6793 S E5, E4+NT
                 E MUSCLE RELAXANT/CT
L45
            5669 S E4-E10
                 E E4+ALL
L46
            8857 S E5, E6, E4+NT
                 E DECONGESTANT/CT
L47
             431 S E4, E5
                 E E4+ALL
L48
             431 S E4
                 E BRONCHODIAL/CT
L49
            5474 S E7-E9
                 E E7+ALL
L50
            9708 S E5, E4+NT
L51
           66379 S ANTIARTHRIT? OR ANTIHISTAMIN? OR ANTI() (ARTHRIT? OR HISTAMIN?
L52
             147 S L25 AND L42-L51
L53
             120 S L52 AND L39
L54
             13 S L52 AND L41
L55
             13 S L40 AND L54
                 SEL DN AN 3 10 11
L56
               3 S E1-E9
             131 S L41, L52, L53 AND L23
L57
             26 S L57 AND L26
L58
L59
             23 S L58 NOT L56
             15 S L59 NOT L55
L60
                 SEL DN AN 11
L61
               1 S L60 AND E10-E12
L62
               2 S L22
L63
               6 S L56, L61, L62
                 E RAFFA R/AU
            177 S E4-E9
L64
                 E COWAN A/AU
L65
             236 S E3-E15,E17,E20,E21
                 E TALLARIDA R/AU
            103 S E4-E6
L66
L67
               1 S L64-L66 AND L25
L68
               6 S L63, L67 AND L23-L67
                SEL HIT RN
     FILE 'REGISTRY' ENTERED AT 11:00:56 ON 27 JAN 2003
L69
             11 S E1-E11
L70
             10 S L69 NOT C16H25NO2
L71
              1 S L69 NOT L70
=> fil reg
FILE 'REGISTRY' ENTERED AT 11:01:43 ON 27 JAN 2003
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FILE 'REGISTRY' ENTERED AT 11:01:43 ON 27 JAN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ${\tt ZIC/VINITI}$ data file provided by InfoChem.

STRUCTURE FILE UPDATES: 24 JAN 2003 HIGHEST RN 481628-73-3 DICTIONARY FILE UPDATES: 24 JAN 2003 HIGHEST RN 481628-73-3

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

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L70 ANSWER 1 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN **246858-10-6** REGISTRY

CN D-Glucose, 2-amino-2-deoxy-, 3-benzoyl-.alpha.-methylbenzeneacetate (salt) (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Benzeneacetic acid, 3-benzoyl-.alpha.-methyl-, compd. with 2-amino-2-deoxy-D-glucose (1:1) (9CI)

FS STEREOSEARCH

MF C16 H14 O3 . C6 H13 N O5

SR CA

LC STN Files: CA, CAPLUS, USPATFULL

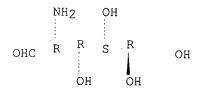
> CM 1

CRN 22071-15-4 CMF C16 H14 O3

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).



1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 131:291311

L70 ANSWER 2 OF 10 REGISTRY COPYRIGHT 2003 ACS

237742-82-4 REGISTRY

D-Glucose, 2-amino-2-deoxy-, (.alpha.R)-3-benzoyl-.alpha.methylbenzeneacetate (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Benzeneacetic acid, 3-benzoyl-.alpha.-methyl-, (.alpha.R)-, compd. with 2-amino-2-deoxy-D-glucose (1:1) (9CI)

FS STEREOSEARCH MF C16 H14 O3 . C6 H13 N O5

SR CA

LC STN Files: CA, CAPLUS, USPATFULL

CM 1

CRN 56105-81-8 CMF C16 H14 O3

Absolute stereochemistry. Rotation (-).

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 131:161659

L70 ANSWER 3 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN **56105-81-8** REGISTRY

OTHER CA INDEX NAMES:

CN Benzeneacetic acid, 3-benzoyl-.alpha.-methyl-, (R)-

OTHER NAMES:

CN (-)-2-(3-Benzoylphenyl)propionic acid

CN (-)-3-Benzoyl-.alpha.-methylbenzeneacetic acid

CN (-)-Ketoprofen

CN (2R)-2-(3-Benzoylphenyl)propionic acid

CN (R)-(-)-Ketoprofen

CN (R)-2-(3-Benzoylphenyl)propionic acid

CN (R)-3-Benzoyl-.alpha.-methylphenylacetic acid

CN (R)-Ketoprofen

FS STEREOSEARCH

MF C16 H14 O3

CI COM

LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAPLUS, CASREACT, CEN, CHEMINFORMRX, CIN, DRUGNL, DRUGPAT, DRUGUPDATES, IPA, PHAR, PROMT, TOXCENTER, USPAT2, USPATFULL (*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

307 REFERENCES IN FILE CA (1962 TO DATE)
8 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
309 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:11225
REFERENCE 2: 137:375140
...
REFERENCE 3: 137:261988

REFERENCE 4: 137:257005

REFERENCE 5: 137:241443

REFERENCE 6: 137:169291

REFERENCE 7: 137:162942

REFERENCE 8: 137:124232

REFERENCE 9: 137:83751

REFERENCE 10: 137:68303

L70 ANSWER 4 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN **51146-56-6** REGISTRY

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)-, (.alpha.S)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)-, (S)-OTHER NAMES:

(+)-(S)-Ibuprofen

CN (+)-(S)-p-Isobutylhydratropic acid

CN (+)-.alpha.-Methyl-4-(2-methylpropyl)benzeneacetic acid

CN (+)-Ibuprofen

CN (+)-Ibuprophen

CN (S)-(+)-2-(4-Isobutylphenyl) propionic acid

CN (S)-(+)-4-Isobutyl-.alpha.-methylphenylacetic acid

CN (S)-(+)-Ibuprofen

CN (S)-2-(4-Isobutylphenyl)propanoic acid

CN (S)-2-(4-Isobutylphenyl)propionic acid

CN (S)-2-(p-Isobutylphenyl)propionic acid

CN (S)-Ibuprofen

CN d-Ibuprofen

CN Dexibuprofen

CN Seractil

FS STEREOSEARCH

MF C13 H18 O2

CI COM

LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*,
BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, CBNB, CEN,
CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, DRUGPAT, EMBASE, IPA,
MEDLINE, PHAR, PROMT, SYNTHLINE, TOXCENTER, USAN, USPAT2, USPATFULL
(*File contains numerically searchable property data)
Other Sources: WHO

Absolute stereochemistry. Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

762 REFERENCES IN FILE CA (1962 TO DATE)

17 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

762 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:65749

REFERENCE 2: 138:16628

REFERENCE 3: 138:16626

REFERENCE 4: 138:16625

REFERENCE 5: 138:8366

REFERENCE 6: 137:369829

REFERENCE 7: 137:369804

REFERENCE 8: 137:369721

REFERENCE 9: 137:359575

REFERENCE 10: 137:299925

L70 ANSWER 5 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 29031-19-4 REGISTRY

CN D-Glucose, 2-amino-2-deoxy-, sulfate (salt) (8CI, 9CI) (CA INDEX NAME) OTHER NAMES:

CN D-Glucosamine sulfate

CN DONA

CN Glucosamine sulfate

FS STEREOSEARCH

DR 216447-61-9

MF C6 H13 N O5 . x H2 O4 S

CI COM

LC STN Files: ADISINSIGHT, ADISNEWS, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAPLUS, CHEMCATS, CHEMLIST, CIN, CSCHEM, DIOGENES, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, PHAR, PROMT, TOXCENTER, USPAT2, USPATFULL

(*File contains numerically searchable property data) Other Sources: EINECS**

(**Enter CHEMLIST File for up-to-date regulatory information)

CM 1

CRN 7664-93-9 CMF H2 O4 S

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

123 REFERENCES IN FILE CA (1962 TO DATE)
4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
125 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:32807

REFERENCE 2: 138:8364

REFERENCE 3: 137:358197

REFERENCE 4: 137:345779

REFERENCE 5: 137:329479

REFERENCE 6: 137:268432

REFERENCE 7: 137:241878

REFERENCE 8: 137:222118

REFERENCE 9: 137:222055

REFERENCE 10: 137:210995

L70 ANSWER 6 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN **22071-15-4** REGISTRY

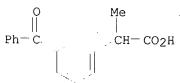
CN Benzeneacetic acid, 3-benzoyl-.alpha.-methyl- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Hydratropic acid, m-benzoyl- (8CI)

OTHER NAMES:

```
(.+-.)-2-(3-Benzoylphenyl)propionic acid
CN
     (.+-.)-3-Benzoyl-.alpha.-methylbenzeneacetic acid
CN
     (.+-.)-Ketoprofen
CN
     (.+-.)-m-Benzoylhydratropic acid
CN
CN
     (RS)-Ketoprofen
CN
     .alpha.-(3-Benzoylphenyl)propionic acid
CN
     2-(3-Benzoylphenyl)propionic acid
CN
CN
     2-(m-Benzoylphenyl)propionic acid
     3-Benzoyl-.alpha.-methylbenzeneacetic acid
CN
CN
     3-Benzoylhydratropic acid
CN
     Alrheumun
CN
     Aneol
CN
     Bi-profenid
CN
     Capisten
CN
     Epatec
CN
     Ketoprofen
CN
     Ketoprofene
CN
     Ketoprophen
CN
     Ketorin
CN
    m-Benzoylhydratropic acid
CN
     Orudis
CN
     Oruvail
CN
     Profenid
CN
     R.P. 19583
     Racemic ketoprofen
CN
CN
     RU 4733
FS
     3D CONCORD
DR
     172964-50-0, 22161-86-0
MF
     C16 H14 O3
CI
LC
     STN Files:
                  ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
       BIOTECHNO, CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
      CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DETHERM*, DIOGENES,
      DRUGPAT, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*,
      MSDS-OHS, NIOSHTIC, PHAR, PHARMASEARCH, PROMT, RTECS*, SPECINFO,
      SYNTHLINE, TOXCENTER, ULIDAT, USAN, USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
    Other Sources:
                     EINECS**, NDSL**, TSCA**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
```



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2897 REFERENCES IN FILE CA (1962 TO DATE)
90 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
2906 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:61373

REFERENCE 2: 138:61369

REFERENCE 3: 138:60879

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REFERENCE
            4:
                138:49359
REFERENCE
                138:44736
            5:
REFERENCE
            6:
                138:37373
                138:33289
REFERENCE
            7:
                138:32500
REFERENCE
            8:
REFERENCE
                138:16628
            9:
REFERENCE 10:
                138:16626
     ANSWER 7 OF 10 REGISTRY COPYRIGHT 2003 ACS
RN
     15687-27-1 REGISTRY
CN
     Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI)
                                                                      (CA INDEX
     NAME)
OTHER NAMES:
     (.+-.)-.alpha.-Methyl-4-(2-methylpropyl)benzeneacetic acid
     (.+-.)-2-(p-Isobutylphenyl) propionic acid
CN
CN
     (.+-.)-Ibuprofen
     (.+-.)-Ibuprophen
CN
CN
     (4-Isobutylphenyl) -. alpha. -methylacetic acid
CN
     (RS)-Ibuprofen
CN
     (S)-4-Isobutyl-.alpha.-methylphenylacetic acid
CN
     .alpha.-(4-Isobutylphenyl)propionic acid
CN
     .alpha.-Methyl-4-(2-methylpropyl)benzeneacetic acid
CN
     2-(4'-Isobutylphenyl)propionic acid
CN
     2-(4-Isobutylphenyl)propanoic acid
CN
     2-(p-Isobutylphenyl)propionic acid
CN
     4-Isobutyl-.alpha.-methylphenylacetic acid
CN
     4-Isobutylhydratropic acid
CN
     Act 3
CN
     Adex 200
CN
     Advil
CN
     Alaxan
CN
     Algofen
CN
     Am-Fam 400
     Anafen
CN
CN
     Anco
CN
     Andran
CN
     Anflagen
CN
     Antarene
CN
     Antiflam
CN
     Apo-Ibuprofen
CN
     Apsifen
CN
     Artofen
CN
     Artril
CN
     Atril 300
CN
     Balkaprofen
CN
     Betaprofen
CN
     Bloom
CN
     Bluton
CN
     Brofen
CN
     Brufanic
CN
     Brufen
CN
     Brufen 400
CN
     Brufen Retard
CN
     Bruflam
CN
     Brufort
CN
     Buburone
```

CN

Burana

```
CN
    Butacortelone
    Carol
CN
CN
    Cobo
CN
    Codral Period Pain
CN
     Combiflam
    Dibufen
CN
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
     DISPLAY
FS
     3D CONCORD
DR
     58560-75-1, 139466-08-3
MF
    C13 H18 O2
CI
    COM
LC
    STN Files:
                 ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*,
       BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT,
       CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, DDFU,
       DIOGENES, DIPPR*, DRUGPAT, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT, IFIUDB,
       IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC, PHAR, PHARMASEARCH, PIRA,
       PROMT, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, ULIDAT, USAN, USPAT2,
       USPATFULL, VETU
         (*File contains numerically searchable property data)
                      DSL**, EINECS**, TSCA**, WHO
    Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
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PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

5852 REFERENCES IN FILE CA (1962 TO DATE)
174 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
5866 REFERENCES IN FILE CAPLUS (1962 TO DATE)
2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 138:65749
REFERENCE 2: 138:61373

REFERENCE 3: 138:61369

REFERENCE 4: 138:61365

REFERENCE 5: 138:61315

REFERENCE 6: 138:61149

REFERENCE 7: 138:60879

REFERENCE 8: 138:55713

REFERENCE 9: 138:44736

REFERENCE 10: 138:44722

L70 ANSWER 8 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN **7512-17-6** REGISTRY

D-Glucose, 2-(acetylamino)-2-deoxy- (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES: D-Glucose, 2-acetamido-2-deoxy- (8CI) OTHER NAMES: 2-Acetamido-2-deoxy-D-glucose CN 2-Acetamido-2-deoxyglucose CN CN 2-Acetamido-D-glucose 2-Acetylamino-2-deoxy-D-glucose CN Acetylglucosamine CN D-N-Acetylglucosamine CN Marine Sweet CN CN N-Acetyl-2-amino-2-deoxy-D-glucose N-Acetyl-2-amino-2-deoxyglucose CN CN N-Acetyl-D-glucosamine CN N-Acetylglucosamine FS STEREOSEARCH 7132-76-5, 134-61-2, 173382-53-1, 98632-70-3 DR MF C8 H15 N O6 CI COM ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, LC STN Files: BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMLIST, CIN, CSCHEM, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT, PIRA, PROMT, SPECINFO, TOXCENTER, USPAT2, USPATFULL (*File contains numerically searchable property data) Other Sources: DSL**, EINECS**, TSCA** (**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

4975 REFERENCES IN FILE CA (1962 TO DATE)
372 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
4980 REFERENCES IN FILE CAPLUS (1962 TO DATE)
1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 138:61045 REFERENCE 2: 138:51620

REFERENCE 3: 138:51537

REFERENCE 4: 138:44821

REFERENCE 5: 138:37402

REFERENCE 6: 138:36689

REFERENCE 7: 138:34830

REFERENCE 8: 138:23767

REFERENCE 9: 138:21926

REFERENCE 10: 138:21284

L70 ANSWER 9 OF 10 REGISTRY COPYRIGHT 2003 ACS

3416-24-8 REGISTRY

CN D-Glucose, 2-amino-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

OTHER NAMES:

2-Amino-2-deoxy-D-glucopyranose

2-Amino-2-deoxy-D-glucose

CN 2-Amino-2-deoxyglucose

CN 2-Deoxy-2-amino-D-glucose

CN 2-Deoxy-2-aminoglucose

CN Chitosamine

CN D-Glucosamine

CN Glucosamine

FS STEREOSEARCH

DR 58-87-7, 58267-75-7, 2351-15-7

MF C6 H13 N O5

CI

LC ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, STN Files: BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMLIST, CIN, CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, HODOC*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT, NIOSHTIC, PHARMASEARCH, PIRA, PROMT, RTECS*, SYNTHLINE, TOXCENTER, TULSA, USAN, USPAT2, USPATFULL, VETU (*File contains numerically searchable property data) Other Sources:

EINECS**, NDSL**, TSCA**, WHO (**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

4730 REFERENCES IN FILE CA (1962 TO DATE)

312 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

4733 REFERENCES IN FILE CAPLUS (1962 TO DATE) 6 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 138:61295

REFERENCE 2: 138:61290

REFERENCE 3: 138:52546

REFERENCE 138:49392 4:

REFERENCE 5: 138:44821

REFERENCE 6: 138:44738

REFERENCE 7: 138:44521 REFERENCE 8: 138:44100

REFERENCE 9: 138:37189

REFERENCE 10: 138:35679

L70 ANSWER 10 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 66-84-2 REGISTRY

CN D-Glucose, 2-amino-2-deoxy-, hydrochloride (8CI, 9CI) (CA INDEX NAME)

OTHER NAMES:

CN 2-Amino-2-deoxy-D-glucose hydrochloride

CN 2-Deoxy-2-amino-D-glucose hydrochloride

CN Chitosamine hydrochloride

CN Cosamin

CN D-(+)-Glucosamine hydrochloride

CN D-Glucosamine chloride

CN D-Glucosamine hydrochloride

CN Glucosamine hydrochloride

FS STEREOSEARCH

DR 2002-25-7, 3615-52-9, 66573-21-5, 151799-45-0, 34673-29-5, 214046-22-7

MF C6 H13 N O5 . Cl H

CI COM

LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAOLD, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMLIST, CIN, CSCHEM, IFICDB, IFIPAT, IFIUDB, IPA, PIRA, PROMT, RTECS*, TOXCENTER, ULIDAT, USPAT2, USPATFULL

(*File contains numerically searchable property data)

Other Sources: DSL**, EINECS**, TSCA**

(**Enter CHEMLIST File for up-to-date regulatory information)

CRN (3416-24-8)

Absolute stereochemistry. Rotation (+).

● HCl

769 REFERENCES IN FILE CA (1962 TO DATE)

17 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

772 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:61045

REFERENCE 2: 138:51537

REFERENCE 3: 138:49392

REFERENCE 4: 138:39499

REFERENCE 5: 138:14820

REFERENCE 6: 138:11371

REFERENCE 7: 138:8378

REFERENCE 8: 138:8364

REFERENCE 9: 138:4803

REFERENCE 10: 137:386237

=> d 171 ide can

L71 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS

RN **27203-92-5** REGISTRY

CN Cyclohexanol, 2-[(dimethylamino)methyl]-1-(3-methoxyphenyl)-, (1R,2R)-rel-(9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Cyclohexanol, 2-[(dimethylamino)methyl]-1-(3-methoxyphenyl)-, cis-(.+-.)-

CN Cyclohexanol, 2-[(dimethylamino)methyl]-1-(m-methoxyphenyl)- (8CI)

OTHER NAMES:

CN (.+-.)-Tramadol

CN cis-Tramadol

CN Racemic tramadol

CN Tramadol

FS STEREOSEARCH

DR 113683-92-4, 73806-46-9

MF C16 H25 N O2

CI CON

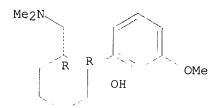
LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*,
BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CBNB,
CEN, CHEMCATS, CHEMLIST, CIN, DDFU, DIOGENES, DRUGU, EMBASE, IPA,
MEDLINE, MRCK*, PHAR, PHARMASEARCH, PROMT, RTECS*, SYNTHLINE, TOXCENTER,
USAN, USPAT2, USPATFULL, VETU

(*File contains numerically searchable property data)

Other Sources: EINECS**, WHO

(**Enter CHEMLIST File for up-to-date regulatory information)

Relative stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

532 REFERENCES IN FILE CA (1962 TO DATE)

18 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

537 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:61448

REFERENCE 2: 138:49342

REFERENCE 3: 138:33355

REFERENCE 4: 138:33351

REFERENCE 138:32766 5:

REFERENCE 138:19392 6:

REFERENCE 7: 137:389167

REFERENCE 8: 137:375269

REFERENCE 9: 137:363089

REFERENCE 10: 137:363086

=> fil hcaplus

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FILE COVERS 1907 - 27 Jan 2003 VOL 138 ISS 5 FILE LAST UPDATED: 26 Jan 2003 (20030126/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d 168 all hitstr tot

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L68 ANSWER 1 OF 6 HCAPLUS COPYRIGHT 2003 ACS
```

AN 2002:256052 HCAPLUS

DN 136:284456

TΙ Analgesic and glucosamine compositions

ΙN Raffa, Robert; Cowan, Alan; Tallarida, Ronald

Temple University of the Commonwealth System of Higher Education, USA PΑ

SO PCT Int. Appl., 24 pp.

CODEN: PIXXD2

DT Patent

English LΑ

IC ICM A61K031-70

CC 63-6 (Pharmaceuticals)

FAN.	CNT	1																
	PA	rent	NO.		KI	ND	DATE			A	PPLI	CATI	ON NO	٥.	DATE			
										_		- -						
ΡI	WO	2002	0262	39	A	1	2002	0404		W	0 2 C	01-U	S296	06	2001	0921		
		₩:	ΑE,	AG,	ΑL,	ΑM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	ΕE,	ES,	FI,	GB,	GD,	GE,	GH,
			GM,	HR,	ΗU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PH,	PL,
			PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,
			UZ,	VN,	YU,	ZA,	ZW,	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM		

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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
              DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
      AU 2001092929
                        A5
                              20020408
                                             AU 2001-92929
                                                               20010921
     US 2002058642
                        Α1
                              20020516
                                             US 2001-964178
                                                               20010925
PRAI US 2000-235405P
                              20000926
     WO 2001-US29606
                        W
                              20010921
     This invention relates to a compn. a glucosamine material and an
AB
     analgesic compd. such as a nonsteroidal anti-
     inflammatory drug (NSAID) and/or an opioid
     analgesic and its use for treatment of pain in pharmaceutical or
     veterinary applications. When the components are administered within
     certain ratios, the analgesic efficacy of the compn. is
     super-additive (synergistic) relative to the analgesic efficacy
     of the analgesic compd. alone. Solns. of glucosamine
     with ibuprofen or ketoprofen were given.
ST
     analgesic NSAID glucosamine compn
ΙT
     Analgesics
       Antiarthritics
       Antihistamines
       Bronchodilators
       Decongestants
       Muscle relaxants
         (analgesic and glucosamine compns.)
ΙT
     Anti-inflammatory agents
         (nonsteroidal; analgesic and glucosamine
        compns.)
ΙT
     Drug interactions
         (synergistic; analgesic and glucosamine compns.)
     66-84-2, Glucosamine hydrochloride
     Acetaminophen 3416-24-8, Glucosamine 7512-17-6
     , N-Acetylglucosamine 15307-86-5, Diclofenac
     15687-27-1, Ibuprofen 22071-15-4,
     Ketoprofen 27203-92-5, Tramadol 29031-19-4,
     Glucosamine sulfate
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (analgesic and glucosamine compns.)
RE.CNT
              THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Armitage; US 4501727 A 1985 HCAPLUS
     66-84-2, Glucosamine hydrochloride 3416-24-8,
IT
     Glucosamine 7512-17-6, N-Acetylglucosamine
     15687-27-1, Ibuprofen 22071-15-4,
     Ketoprofen 27203-92-5, Tramadol 29031-19-4,
     Glucosamine sulfate
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (analgesic and glucosamine compns.)
RN
     66-84-2 HCAPLUS
CN
     D-Glucose, 2-amino-2-deoxy-, hydrochloride (8CI, 9CI) (CA INDEX NAME)
Absolute stereochemistry. Rotation (+).
```

RN 3416-24-8 HCAPLUS

CN D-Glucose, 2-amino-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 7512-17-6 HCAPLUS

CN D-Glucose, 2-(acetylamino)-2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 15687-27-1 HCAPLUS

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI) (CA INDEX NAME)

RN 22071-15-4 HCAPLUS

CN Benzeneacetic acid, 3-benzoyl-.alpha.-methyl- (9CI) (CA INDEX NAME)

RN 27203-92-5 HCAPLUS

CN Cyclohexanol, 2-[(dimethylamino)methyl]-1-(3-methoxyphenyl)-, (1R,2R)-rel-(9CI) (CA INDEX NAME)

Relative stereochemistry.

CM 1

CRN 7664-93-9 CMF H2 O4 S

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

L68 ANSWER 2 OF 6 HCAPLUS COPYRIGHT 2003 ACS

AN 1999:672575 HCAPLUS

DN 131:291311

TI Pharmaceutical preparations containing hydrosoluble **ketoprofen** salts

IN Giorgetti, Paolo Luca Maria

PA Errekappa Euroterapici S.p.A., Italy

SO PCT Int. Appl., 36 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K031-40 ICS A61K031-70; A61K031-19; C07H005-06; C07D207-16; C07C051-41; C07C059-84

CC 63-6 (Pharmaceuticals)

FAN.CNT 1

r AN.	CNII						
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
ΡI	WO 9952528	A1	19991021	WO 1999-IB626	19990409		

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W: CA, CN, JP, KR, US
          RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
              PT, SE
      CH 692939
                        Д
                              20021231
                                             CH 1998-843
                                                              19980411
      EP 1024808
                        A1
                             20000809
                                             EP 1999-910606
                                                              19990409
              AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, FI
      JP 2002510336
                                             JP 1999-551422
                        T2
                             20020402
                                                              19990409
      US 6291527
                                             US 1999-445672
                        В1
                             20010918
                                                              19991210
PRAI CH 1998-843
                             19980411
                        Α
     CH 1999-618
                        Α
                             19990331
      WO 1999-IB626
                        W
                             19990409
AΒ
     The new pharmaceutical prepns. contain hydrosol. salts obtained through a
     reaction between {\bf ketoprofen} and {\bf glucosamine} and/or
     Proline and/or Hydroxyproline from 0.01 to 30 % of the mass.
     are useful for anti-inflammatory and {\bf analgesic} treatment of
     joints and mucous membranes. Thus, an i.m. injection (quantities for 1
     unit) consisted of ketoprofen glucosamine salt 170
     equiv. to ketoprofen acid 100 mg, excipients such as benzyl alc.
     90, NaCl 27 mg and water for injectable prepns. up to 3 mL.
ST
     ketoprofen salt pharmaceutical hydrosoluble
ΙT
     Drug delivery systems
         (capsules, controlled-release; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
IT
     Drug delivery systems
         (capsules; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
IT
     Drug delivery systems
         (chewing gums; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
ΙT
     Drug delivery systems
         (foams; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
ΙT
     Drug delivery systems
         (gels; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
ΙT
     Drug delivery systems
         (granules, pharmaceutical; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
ΙT
     Drug delivery systems
        (granules, sustained release; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
ΙT
     Drug delivery systems
        (injections, i.m.; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
IΤ
     Drug delivery systems
        (injections, i.v.; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
IT
     Drug delivery systems
        (injections; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
IT
     Drug delivery systems
        (lotions; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
IΤ
     Drug delivery systems
        (ointments, creams; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
ΙT
     Drug delivery systems
        (ointments; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
IT
     Drug delivery systems
        (oral; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
```

```
Analgesics
       Anti-inflammatory agents
     Mouthwashes
         (pharmaceutical prepns. contg. hydrosol. ketoprofen salts)
ΙT
     Drug delivery systems
         (solns., ophthalmic; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
ΙT
     Drug delivery systems
         (solns.; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
ΙT
     Drug delivery systems
         (suppositories; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
TT
     Drug delivery systems
        (suspensions, sustained-release; pharmaceutical prepns. contq.
        hydrosol. ketoprofen salts)
TΤ
     Drug delivery systems
        (suspensions; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
TΤ
     Drug delivery systems
        (tablets, coated; pharmaceutical prepns. contg. hydrosol.
        ketoprofen salts)
ΙT
     Drug delivery systems
        (tablets; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
ΙT
     Drug delivery systems
        (topical; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
ΙT
     Drug delivery systems
        (transdermal; pharmaceutical prepns. contq. hydrosol.
        ketoprofen salts)
ΙT
     Drug delivery systems
        (vaginal; pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
IΤ
     246858-10-6
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (9099000 pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
ΙT
     22071-15-4D, Ketoprofen, salts
                                       246858-11-7
     246858-12-8
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pharmaceutical prepns. contg. hydrosol. ketoprofen salts)
              THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Green Cross Corporation; JP 63093718 A 1988 HCAPLUS
(2) Hoechst Celanese Corporation; WO 9412451 A 1994 HCAPLUS
(3) Laboratorios Menarini; WO 9616016 A 1996 HCAPLUS
(4) The Procter & Gamble Company; WO 9507079 A 1995 HCAPLUS
(5) Veronesi; US 4748174 A 1988 HCAPLUS
ΙT
     246858-10-6
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (9099000 pharmaceutical prepns. contg. hydrosol. ketoprofen
        salts)
RN
     246858-10-6 HCAPLUS
CN
     D-Glucose, 2-amino-2-deoxy-, 3-benzoyl-.alpha.-methylbenzeneacetate (salt)
     (9CI) (CA INDEX NAME)
     CM
          1
     CRN 22071-15-4
```

CMF C16 H14 O3

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

IT 22071-15-4D, Ketoprofen, salts

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceutical prepns. contg. hydrosol. ketoprofen salts)

RN 22071-15-4 HCAPLUS

CN Benzeneacetic acid, 3-benzoyl-.alpha.-methyl- (9CI) (CA INDEX NAME)

L68 ANSWER 3 OF 6 HCAPLUS COPYRIGHT 2003 ACS

AN 1999:551706 HCAPLUS

DN 131:161659

Pharmaceutical preparations containing (R)-2-(3-Benzoylphenyl)propionic acid salts for the treatment of neutrophil-dependent diseases and phlogistic processes

IN Bertini, Riccardo; Bizzarri, Cinzia; Brandolini, Laura; Melillo, Gabriella; Caselli, Gianfranco; Clavenna, Gaetano

PA Dompe S.p.A., Italy

SO Eur. Pat. Appl., 11 pp.

CODEN: EPXXDW

DT Patent

LA English

IC ICM A61K031-19

CC 63-6 (Pharmaceuticals)

Section cross-reference(s): 1

FAN.CNT 1

	PA:	rent	NO.		KI	ND	DATE			А	PPLI	CATI	ON NO	ο.	DATE			
PI		9359 9359			— <u>—</u> А	_	1999			E	P 19	99-1	0132	2	1999	0125		
		R:	AT, IE,	BE, SI,	CH, LT,	DE, LV,	DK, FI,	ES, RO	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,

```
IT 1298214
                        В1
                             19991220
                                            IT 1998-MI146
                                                             19980128
     US 6069172
                       Α
                             20000530
                                            US 1999-237901
                                                             19990127
PRAI IT 1998-MI146
                       Α
                             19980128
     A new use of the enantiomer (R)-ketoprofen and of its salts with
     suitable org. and inorg. bases in the therapy of neutrophil-dependent
     diseases and phlogistic processes is described as well as pharmaceutical
     prepns. contg. such compds. and useful for oral, parenteral or topical
     administration. Specific inhibitory effects of lysine salts of (R)- and
     (S)-ketoprofen on interleukin-8- stimulated chemotactic
     migration are shown. The effects of these drugs were not limited to
     interleukin-8-stimulated chemotaxis, but were also displayed,
     surprisingly, on the processes induced by other physiol and non-physiol.
     stimulant acting, although in different ways, through variations in
     intracellular Ca++ concn. An injection soln. contained (R)-
     ketoprofen lysine salt 80, citric acid 2.5, sodium hydrate 1.5 mg,
     and water 1 mL.
     pharmaceutical enantiomer ketoprofen salt neutrophil disease;
     injection pharmaceutical lysine ketoprofen
IT
     Neutrophil
        (-dependent diseases; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
IT
     Reperfusion
        (damage from; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
IΤ
     Drug delivery systems
        (emulsions; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
IT
     Lung, disease
        (fibrosis, idiopathic; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (foams; pharmaceutical prepns. contg. enantiomer (R)-ketoprofen
        and of its salts for treatment of neutrophil-dependent diseases and
        phlogistic processes)
ΙT
     Kidney, disease
        (glomerulonephritis; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (granules; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (injections; pharmaceutical prepns. contq. enantiomer (R)-
       ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (ointments, creams; pharmaceutical prepns. contq. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (ointments; pharmaceutical prepns. contg. enantiomer (R)-
       ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (parenterals; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
       neutrophil-dependent diseases and phlogistic processes)
ΙT
     Psoriasis
```

```
(pharmaceutical prepns. contg. enantiomer (R)-ketoprofen and
        of its salts for treatment of neutrophil-dependent diseases and
        phlogistic processes)
TΤ
     Drug delivery systems
         (powders; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
TΤ
     Breathing (animal)
         (respiratory failure, acute; pharmaceutical prepns. contg. enantiomer
         (R)-ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
         (solns.; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (sprays; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (suppositories; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (tablets; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     Drug delivery systems
        (topical; pharmaceutical prepns. contg. enantiomer (R)-
        ketoprofen and of its salts for treatment of
        neutrophil-dependent diseases and phlogistic processes)
ΙT
     56105-81-8D, salts
                          167300-65-4
                                        167300-66-5
                                                       167300-67-6
     167300-68-7
                   167300-69-8
                                 167300-70-1
                                                237742-71-1
                                                              237742-72-2
     237742-73-3
                   237742-74-4
                                 237742-75-5
                                                237742-76-6
                                                              237742-77-7
     237742-78-8
                   237742-79-9
                                 237742-80-2
                                               237742-81-3 237742-82-4
     237742-83-5
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (pharmaceutical prepns. contg. enantiomer (R)-ketoprofen and
        of its salts for treatment of neutrophil-dependent diseases and
        phlogistic processes)
ΤТ
     56105-81-8D, salts 237742-82-4
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (pharmaceutical prepns. contg. enantiomer (R)-ketoprofen and
        of its salts for treatment of neutrophil-dependent diseases and
        phlogistic processes)
RN
     56105-81-8 HCAPLUS
CN
     Benzeneacetic acid, 3-benzoyl-.alpha.-methyl-, (.alpha.R)- (9CI) (CA
     INDEX NAME)
Absolute stereochemistry. Rotation (-).
```



RN 237742-82-4 HCAPLUS

D-Glucose, 2-amino-2-deoxy-, (.alpha.R)-3-benzoyl-.alpha.-CN methylbenzeneacetate (9CI) (CA INDEX NAME)

CM1

CRN 56105-81-8 CMF C16 H14 O3

Absolute stereochemistry. Rotation (-).

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

L68 ANSWER 4 OF 6 HCAPLUS COPYRIGHT 2003 ACS

AN 1997:224073 HCAPLUS

DN 126:216664

TΙ Pharmaceutical compositions containing analgesics and antihistamines and methods for treating respiratory disorders
Cramer, Ronald Dean; Mitra, Sekhar; Riker, Donald Kay

ΙN

PΑ Procter and Gamble Company, USA

SO PCT Int. Appl., 19 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K045-06

ICS A61K031-40; A61K031-19

ICI A61K031-40, A61K031-19; A61K031-40, A61K031-19, A61K031-135; A61K031-40, A61K031-19, A61K031-485

63-6 (Pharmaceuticals)

FAN.CNT 1

	PA?	rent	NO.		KI	ND .	DATE			A	PPLI	CATI	ON N	0.	DATE				
										_					 -				
ΡI	WO	9704	808		Α	1	1997	0213		M	0 19	96 - U	S122	49	1996	0725			
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																	LK,		
			LS,	LT,	LU,	LV,	MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	
			SD,										,		•	•	•	•	
		RW:	KE,	LS,	MW,	SD,	SZ,	UG,	AT,	BE,	CH,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	

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IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA
      CA 2227958
                        AA
                            19970213
                                           CA 1996-2227958 19960725
      AU 9665991
                        A1
                             19970226
                                            AU 1996-65991
                                                             19960725
      EP 841947
                        Α1
                             19980520
                                            EP 1996-925495
                                                             19960725
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, FI
      JP 11510168
                                           JP 1996-507747 19960725
                       T2
                           19990907
      ZA 9606385
                                           ZA 1996-6385
                        Α
                             19970604
                                                             19960728
 PRAI US 1995-508775
                             19950728
                       Α
      US 1996-611528
                       Α
                             19960305
     WO 1996-US12249 W
                             19960725
 OS
     MARPAT 126:216664
     Compns. and methods for providing improved treatment, management or
     mitigation of cold cold-like, allergy, sinus and/or flu symptoms by
     administering a safe and effective amt. of a compn. comprising an
     analgesic agent along with certain pyrrolidine and piperidine
     ether antihistaminic agents. A hard gelatin capsule contained
     ibuprofen 200.00, clemastine fumarate 0.67, pseudoephedrine. HCl
     30.00 mg, and lactose q.s. Administration of 1-2 capsules every 4-12\ h
     provide relief from cough, cold, flu and allergic rhinitis symptoms.
     pharmaceutical analgesic antihistamine respiratory
     disorder; capsule clemastine pseudoephedrine cold flu cough
     Drug delivery systems
         (capsules; pharmaceutical compns. contg. analgesics and
        antihistamines for treating respiratory disorders)
IT
     Respiratory tract
        (disease; pharmaceutical compns. contg. analgesics and
        antihistamines for treating respiratory disorders)
ΙT
     Drug delivery systems
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (liqs.; pharmaceutical compns. contg. analgesics and
        antihistamines for treating respiratory disorders)
ΙT
     Allergy inhibitors
       Analgesics
       Anti-inflammatory agents
       Antihistamines
     Antitussives
       Decongestants
     Expectorants
     Influenza
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (pharmaceutical compns. contg. analgesics and
        antihistamines for treating respiratory disorders)
ΙT
    Amino acids, biological studies
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (salts; pharmaceutical compns. contg. analgesics and
        antihistamines for treating respiratory disorders)
ΙT
     Drug delivery systems
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (tablets; pharmaceutical compns. contg. analgesics and
        antihistamines for treating respiratory disorders)
IT
    Adrenoceptor agonists
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
```

(.alpha.-; pharmaceutical compns. contg. analgesics and

```
antihistamines for treating respiratory disorders)
      56-87-1, Lysine, biological studies 58-08-2, Caffeine, biological
TΤ
      studies
                 58-73-1, Diphenhydramine
                                                59-33-6 59-42-7, Phenylephrine
      59-46-1, Procaine 62-49-7, Choline 70-26-8, Ornithine 71-00-1,
      Histidine, biological studies
                                          74-79-3, Arginine, biological studies
      76-57-3, Codeine 77-22-5, Caramiphen 77-23-6, Carbetapentane
      79-09-4D, Propanoic acid, derivs., biological studies 83-67-0,
      Theobromine
                      86-22-6, Brompheniramine 90-82-4, Pseudoephedrine
      91-81-6, Tripelennamine 93-14-1, Glyceryl guaiacolate 100-37-8,
      2-Diethylaminoethanol 102-69-2, Tripropylamine 107-15-3,
      1,2-Ethanediamine, biological studies 107-43-7, Betaine
      110-85-0, Piperazine, biological studies 110-89-4, Piperidine,
      biological studies
                              113-92-8
                                          118-23-0, Bromodiphenhydramine
      Purine
               121-44-8, biological studies
                                                   125-29-1, Hydrocodone
                                                                              125-69-9,
      Dextromethorphan hydrobromide 125-71-3, Dextromethorphan 125-92-8,
                    128-62-1, Noscapine 129-03-3, Cyproheptadine 132-21-8,
      Hydrabamine
      Dexbrompheniramine
                              299-42-3, Ephedrine
                                                      345-78-8, Pseudoephedrine
                        466-99-9, Hydromorphone
      hydrochloride
                                                      486-12-4 486-16-8,
      Carbinoxamine
                        562-10-7
                                    569-59-5
                                                 616-91-1, n-Acetylcysteine
      766-09-6, n-Ethylpiperidine 791-35-5, Chlophedianol
                                                                      2451-01-6, Terpin
      hydrate 3416-24-8, Glucosamine
                                          3572-43-8, Bromhexine
      3964-81-6, Azatadine
                               5104-49-4, Flurbiprofen
                                                              6284-40-8,
                                       12125-02-9, Ammonium chloride, biological
      Methylglucamine
                         7723-51-5
                 14838-15-4, Phenylpropanolamine 14976-57-9, Clemastine
      studies
                 15307-86-5, Diclofenac 15687-27-1, Ibuprofen
      fumarate
      18053-31-1, Fominoben
                                18683-91-5, Ambroxol
                                                           21256-18-8, Oxaprozin
     22071-15-4, Ketoprofen 22204-53-1, Naproxen 25523-97-1, Dexchlorpheniramine 26159-34-2, Naproxen sodium 29216-28-2, Mequitazine 29679-58-1, Fenoprofen 31793-07-4, 31842-01-0, Indoprofen 33005-95-7, Tiaprofen 34580-13-7, K 36330-85-5, Fenbufen 39718-89-3, Alminoprofen 40198-53-6, 40828-46-4, Suprofen 50679-08-8, Terfenadine 51234-28-7, B 52549-17-4, Pranoprofen 53716-49-7, Carprofen 53882-12-5, 55843-86-2, Miroprofen 58581-89-8, Azelastine 60607-34-3, 64294-95-7, Setastine 68844-77-9, Paternia le 76201-68-8
                                   29679-58-1, Fenoprofen 31793-07-4, Pirprofen
                                                              34580-13-7, Ketotifen
                                                             40198-53-6, Tioxaprofen
                                                              51234-28-7, Benoxaprofen
                                                              53882-12-5, Lodoxamide
                                                               60607-34-3, Oxatomide
     64294-95-7, Setastine
                                 68844-77-9, Astemizole
                                                              76201-68-8
                                                                            79516-68-0,
                        79712-55-3, Tazifylline 79794-75-5, Loratidine
      Levocabastine
      83881-51-0, Cetirizine
                                  86181-42-2, Temelastine
                                                                87848-99-5, Acrivastine
      90729-43-4, Ebastine
                                96170-72-8
                                             113403-10-4
                                                               115609-60-4, AHR 11325
      162929-63-7
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
      study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
      (Uses)
         (pharmaceutical compns. contg. analgesics and
         antihistamines for treating respiratory disorders)
ΙT
     3416-24-8, Glucosamine 15687-27-1,
     Ibuprofen 22071-15-4, Ketoprofen
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
         (pharmaceutical compns. contg. analgesics and
         antihistamines for treating respiratory disorders)
RN
     3416-24-8 HCAPLUS
```

Absolute stereochemistry. Rotation (+).

D-Glucose, 2-amino-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

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OHC R R S R OH
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RN 15687-27-1 HCAPLUS

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI) (CA INDEX NAME)

RN 22071-15-4 HCAPLUS

CN Benzeneacetic acid, 3-benzoyl-.alpha.-methyl- (9CI) (CA INDEX NAME)

L68 ANSWER 5 OF 6 HCAPLUS COPYRIGHT 2003 ACS

AN 1995:546946 HCAPLUS

DN 122:274115

TI Compositions containing an amino acid salt of a propionic acid nonsteroidal antiinflammatory agent and at least one of a decongestant, an expectorant, an antihistamine, and an antitussive

IN Mitra, Sekhar

PA Procter and Gamble Co., USA

SO PCT Int. Appl., 17 pp. CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K045-06

ICS A61K031-19; A61K031-445; A61K031-485

CC 63-6 (Pharmaceuticals)

FAN.CNT 1

	PATENT NO.	WIND DAME	
	PAIENI NO.	KIND DATE	APPLICATION NO. DATE
DT	HO 0507100		
ΡI	WO 9507103	A1 19950316	1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1
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	RW: AT, BE,	CH, DE, DK, ES,	FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
	CA 2170488	AA 19950316	
	AU 9476040	Al 19950327	
	EP 719156	A1 19960703	
	R: AT, BE,	CH, DE, DK, ES,	FR, GB, GR, IE, IT, LI, LU, NL, PT, SE
	CN 1130354	A 19960904	CN 1994-193312 19940824
	BR 9407414	A 19961112	BR 1994-7414 19940824
	JP 09502201	T2 19970304	
PRAI	US 1993-116927		000000 10010021
	WO 1994-US9581	19940824	

AB A method for providing improved treatment, management, or mitigation of cold, coldlike, and/or flu symptoms comprises administering a safe and effective amt. of a compn. comprising certain amino acid salts of propionic acid nonsteroidal antiinflammatory agents

```
along with .gtoreq.1 of a decongestant, expectorant,
     antihistamine, and antitussive. Thus, a hard gelatin capsule
     contained naproxen lysinate 200, pseudoephedrine-HCl 30, astemizole 5, and
     glyceryl guaiacolate 100 mg.
ST
     common cold nonsteroidal antiinflammatory
     decongestant; expectorant nonsteroidal
     antiinflammatory common cold; antihistaminic
     nonsteroidal antiinflammatory common cold; antitussive
     nonsteroidal antiinflammatory common cold
ΤТ
     Antihistaminics
     Antitussives
     Common cold
       Decongestants
     Expectorants
     Influenza
        (common cold treatment with amino acid salt of propionic acid
        nonsteroidal antiinflammatory agent and
        decongestant, expectorant, antihistamine, and/or
        antitussive)
ΙT
     Amino acids, biological studies
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (salts with nonsteroidal antiinflammatory drugs;
        common cold treatment with amino acid salt of propionic acid
        nonsteroidal antiinflammatory agent and
        decongestant, expectorant, antihistamine, and/or
        antitussive)
IT
     Inflammation inhibitors
        (nonsteroidal, amino acid salts; common cold treatment with
        amino acid salt of propionic acid nonsteroidal
        antiinflammatory agent and decongestant, expectorant,
        antihistamine, and/or antitussive)
IT
     56-87-1D, L-Lysine, compds. with nonsteroidal
     antiinflammatory drugs
                            58-08-2D, Caffeine, compds. with
     nonsteroidal antiinflammatory drugs 58-73-1,
     Diphenhydramine
                       59-33-6 59-42-7, Phenylephrine
                                                          59-46-1D, Procaine,
     compds. with nonsteroidal antiinflammatory drugs
     62-49-7D, Choline, compds. with nonsteroidal
    antiinflammatory drugs
                            70-26-8D, L-Ornithine, compds. with
    nonsteroidal antiinflammatory drugs
                                          71-00-1D,
    L-Histidine, compds. with nonsteroidal antiinflammatory
            74-79-3D, L-Arginine, compds. with nonsteroidal
    antiinflammatory drugs
                             76-57-3, Codeine
                                               77-22-5, Caramiphen
    77-23-6, Carbetapentane
                              79-09-4D, Propionic acid, derivs., amino acid
            83-67-0D, Theobromine, compds. with nonsteroidal
    salts
                             86-22-6, Brompheniramine
                                                       90-82-4,
    antiinflammatory drugs
    Pseudoephedrine
                     91-81-6, Tripelennamine
                                               93-14-1, Glyceryl guaiacolate
    100-37-8D, 2-Diethylaminoethanol, compds. with nonsteroidal
    antiinflammatory drugs 102-69-2D, Tripropylamine, compds. with
    nonsteroidal antiinflammatory drugs
                                          107-15-3D,
    1,2-Ethanediamine, compds. with nonsteroidal
    antiinflammatory drugs
                            107-43-7D, Betaine, compds. with
    nonsteroidal antiinflammatory drugs
                                         108-01-0D, compds.
    with nonsteroidal antiinflammatory drugs
                                               110-85-0D,
    Piperazine, compds. with nonsteroidal antiinflammatory
            110-89-4D, Piperidine, compds. with nonsteroidal
                            113-92-8, Chlorpheniramine maleate
    antiinflammatory drugs
    118-23-0, Bromodiphenhydramine
                                    120-73-0D, Purine, compds. with
    nonsteroidal antiinflammatory drugs
                                         121-44-8D, compds.
    with nonsteroidal antiinflammatory drugs
                                               125-29-1,
    Hydrocodone
                 125-69-9, Dextromethorphan hydrobromide
    Dextromethorphan 125-92-8D, Hydrabamine, compds. with
```

nonsteroidal antiinflammatory drugs 128-62-1, Noscapine 129-03-3, Cyproheptadine 132-21-8, Dexbrompheniramine 299-42-3, Ephedrine 345-78-8, Pseudoephedrine hydrochloride 486-12-4, Triprolidine 486-16-8, Carbinoxamine Hydromorphone 616-91-1, N-Acetylcysteine 562-10-7 569-59-5 766-09-6D, N-Ethylpiperidine, compds. with nonsteroidal antiinflammatory drugs 791-35-5, Chlophedianol 2451-01-6, Terpin hydrate 3416-24-8D, Glucosamine, compds. with nonsteroidal antiinflammatory drugs 3572-43-8, 3964-81-6, Azatadine 5104-49-4D, Flurbiprofen, amino acid Bromhexine 6284-40-8D, Methylglucamine, compds. with nonsteroidal antiinflammatory drugs 12125-02-9, Ammonium chloride, biological studies 14838-15-4, Phenylpropanolamine 15687-27-1D, Ibuprofen, amino acid salts 18053-31-1, Fominoben 18683-91-5, Ambroxol 21256-18-8D, Oxaprozin, amino acid salts 22071-15-4D, Ketoprofen, amino acid salts 22204-53-1D, Naproxen, amino acid 25523-97-1, Dexchlorpheniramine 29216-28-2, Mequitazine 31793-07-4D, Pirprofen, amino acid salts 31842-01-0D, Indoprofen, amino acid salts 31879-05-7D, Fenoprofen, amino acid salts 33005-95-7D, Tiaprofen, amino acid salts 34580-13-7, Ketotifen 36330-85-5D, Fenbufen, amino acid salts 39718-89-3D, Alminoprofen, amino acid salts 40198-53-6D, Tioxaprofen, amino acid salts 40828-46-4D, Suprofen, amino acid salts 50679-08-8, Terfenadine 51234-28-7D, Benoxaprofen, amino 52549-17-4D, Pranoprofen, amino acid salts 53716-49-7D, acid salts Carprofen, amino acid salts 53882-12-5, Lodoxamide 55843-86-2D, Miroprofen, amino acid salts 57351-43-6 58581-89-8, Azelastine 60607-34-3, Oxatomide 64294-95-7, Setastine 68844-77-9, Astemizole 79516-68-0, Levocabastine 79712-55-3, Tazifylline 79794-75-5, Loratadine 83881-51-0, Cetirizine 86181-42-87848-99-5, Acrivastine 90729-43-4, Ebastine 83881-51-0, Cetirizine 86181-42-2, Temelastine 113403-10-4 115609-60-4, AHR-11325 136013-66-6 158721-32-5 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(common cold treatment with amino acid salt of propionic acid nonsteroidal antiinflammatory agent and decongestant, expectorant, antihistamine, and/or antitussive)

IT 3416-24-8D, Glucosamine, compds. with nonsteroidal antiinflammatory drugs 15687-27-1D

, Ibuprofen, amino acid salts 22071-15-4D,

Ketoprofen, amino acid salts

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

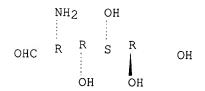
(common cold treatment with amino acid salt of propionic acid nonsteroidal antiinflammatory agent and decongestant, expectorant, antihistamine, and/or antitussive)

3416-24-8 HCAPLUS

RN

CN D-Glucose, 2-amino-2-deoxy- (8CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



RN 15687-27-1 HCAPLUS

Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI) (CA INDEX CN NAME)

22071-15-4 HCAPLUS RN

CN Benzeneacetic acid, 3-benzoyl-.alpha.-methyl- (9CI) (CA INDEX NAME)

L68 ANSWER 6 OF 6 HCAPLUS COPYRIGHT 2003 ACS

ΑN 1992:537660 HCAPLUS

DN 117:137660

ΤI S(+)-phenylalkanoic acids and aminosugar complexes

IN Paradies, Henrich Hasko

MEDICE Chem.-Pharm. Fabrik Puetter GmbH und Co. KG, Germany PΑ

SO Eur. Pat. Appl., 12 pp. CODEN: EPXXDW

DT Patent

LA German

IC ICM C07C057-30

ICS A61K031-205; C07H005-06

CC 63-5 (Pharmaceuticals)

FAN.	CNT	1	·				
	PA'	TENT NO.	KIND	DATE		APPLICATION	NO. DATE
ΡI	ΕP	486046	A2	19920520		EP 1991-119	523 19911115
	EΡ	486046	A3	19921209			
	ΕP	486046	В1	19960501			
		R: AT, BE	, CH, DE	, DK, ES,	FR,	GB, GR, IT, L	I. LU. NL. SE
	HU	59692	A2	19920629			2 19911114
	CA	2055681	AA	19920516		CA 1991-205	
	DE	4137683	A1	19920521		DE 1991-413	
	ΑU	9187904	A1	19920521		AU 1991-879	04 19911115
	ΑU	642309	В2	19931014			
	CN	1061415	A	19920527		CN 1991-110	740 19911115
	zA	9109075	A	19920826		ZA 1991-907	5 19911115
	JΡ	06184003	A2	19940705		JP 1991-354	100 19911115
	JР	2542765	B2	19961009			
	AT	137486	E	19960515		AT 1991-119	523 19911115
	BR	9104997	A	19920623		BR 1991-499	7 19911118
		5604206	A	19970218			722 19940218
PRAI	DE	1990-403646	0	19901115			
	US	1991-792479		19911115			
Λ¢	MAT	רכו 117.12	660				

OS MARPAT 117:137660

AΒ H bridge 1:1 complexes of S-(+)-phenylalkanoic acid drugs with amino sugars are prepd. The complexes have higher bioavailability than the free

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acids. A soln. of 206.27 mg S-(+)-ibuprofen and 236.72 mg
     1-amino-1-deoxy-D-glucitol in 6 mL water was ultrasonicated at 45.degree.,
     to give a 1:1 complex.
ST
     phenylalkanoate amino sugar complex drug
     Drug bioavailability
TΤ
         (of S-(+)-phenylalkanoic acid complexes with amino sugars)
ΙT
     Carbohydrates and Sugars, compounds
     RL: BIOL (Biological study)
         (aminodeoxy, conjugates, with S-(+)-phenylalkanoic acids, for
        high-bioavailability pharmaceuticals)
     488-43-7D, Glucamine, complexes with S-(+)-phenylalkanoic acids
ΤT
     532-19-4D, complexes with S-(+)-phenylalkanoic acids 579-32-8D,
     complexes with S-(+)-phenylalkanoic acids
                                                2494-50-0D, complexes with
     S-(+)-phenylalkanoic acids 3416-24-8D, complexes with
     S-(+)-phenylalkanoic acids
                                  5840-75-5D, complexes with
     S-(+)-phenylalkanoic acids
                                  6284-40-8D, complexes with
     S-(+)-phenylalkanoic acids
                                  6790-34-7D, complexes with
     S-(+)-phenylalkanoic acids
                                  7535-00-4D, complexes with
     S-(+)-phenylalkanoic acids
                                  14216-22-9D, complexes with
     S-(+)-phenylalkanoic acids
                                  14307-02-9D, complexes with
     S-(+)-phenylalkanoic acids
                                  14307-09-6D, complexes with
     S-(+)-phenylalkanoic acids
                                  22204-53-1D, S-(+)-Naproxen, complexes with
     amino sugars
                    26315-48-0D, complexes with S-(+)-phenylalkanoic acids
     27799-64-0D, Allosamine, complexes with S-(+)-phenylalkanoic acids
     51108-70-4D, Ribamine, complexes with S-(+)-phenylalkanoic acids
     51146-56-6D, S-(+)-Ibuprofen, complexes with amino
              83058-22-4D, complexes with S-(+)-phenylalkanoic acids
     RL: BIOL (Biological study)
        (high-bioavailability pharmaceuticals)
IT
     134309-33-4P
                    143381-45-7P
     RL: PREP (Preparation)
        (prepn. of, as high-bioavailability pharmaceutical)
     3416-24-8D, complexes with S-(+)-phenylalkanoic acids
IT
     51146-56-6D, S-(+)-Ibuprofen, complexes with amino
     RL: BIOL (Biological study)
        (high-bioavailability pharmaceuticals)
RN
     3416-24-8 HCAPLUS
     D-Glucose, 2-amino-2-deoxy- (8CI, 9CI) (CA INDEX NAME)
CN
Absolute stereochemistry. Rotation (+).
```

RN 51146-56-6 HCAPLUS
CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)-, (.alpha.S)- (9CI)
(CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

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Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> d ide can tot

L72 ANSWER 1 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 457602-76-5 REGISTRY

CN D-Glucose, 2-(acetylamino)-2-deoxy-, hydrochloride (9CI) (CA INDEX NAME)

FS STEREOSEARCH

MF C8 H15 N O6 . C1 H

SR CA

LC STN Files: CA, CAPLUS

CRN (7512-17-6)

Absolute stereochemistry.

1 REFERENCES IN FILE CA (1962 TO DATE)
1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 137:222087

L72 ANSWER 2 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 127831-02-1 REGISTRY

CN D-Glucose, 2-amino-2-deoxy-, hydrogen sulfate (ester) (9CI) (CA INDEX NAME)

FS STEREOSEARCH

MF C6 H13 N O5 . x H2 O4 S

SR CA

LC STN Files: CA, CAPLUS, USPATFULL

CM 1

CRN 7664-93-9 CMF H2 O4 S

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

2 REFERENCES IN FILE CA (1962 TO DATE)

2 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 136:199305

REFERENCE 2: 113:29136

L72 ANSWER 3 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 51146-57-7 REGISTRY

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)-, (.alpha.R)-(9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)-, (R)-OTHER NAMES:

CN (-)-.alpha.-Methyl-4-(2-methylpropyl)benzeneacetic acid

CN (-)-Ibuprofen

CN (-)-Ibuprophen

CN (R) - (-) - Ibuprofen

CN (R)-2-(4-Isobutylphenyl)propanoic acid

```
CN
     (R)-Ibuprofen
     12: PN: WO02073205 FIGURE: 7 claimed sequence
CN
     1-Ibuprofen
CN
CN
     R-(-)-p-Isobutylhydratropic acid
FS
     STEREOSEARCH
MF
     C13 H18 O2
CI
     COM
LC
     STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
       CA, CAPLUS, CASREACT, CEN, CHEMINFORMRX, CIN, CSCHEM, IPA, PROMT,
       TOXCENTER, USPATFULL
         (*File contains numerically searchable property data)
Absolute stereochemistry. Rotation (-).
                  Bu-i
HO2C.
     Me
```

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

529 REFERENCES IN FILE CA (1962 TO DATE) 3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 529 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:65749 REFERENCE 2: 137:369721

3: 137:242132 REFERENCE

REFERENCE 4: 137:216777

REFERENCE 5: 137:195592

REFERENCE 6: 137:191189

7: 137:150277 REFERENCE

8: 137:149476

REFERENCE 9: 137:114625

REFERENCE 10: 137:83751

L72 ANSWER 4 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 33508-19-9 REGISTRY

D-Glucose, 2-amino-2-deoxy-, sulfate (1:1) (salt) (9CI) (CA INDEX NAME) CN

FS STEREOSEARCH

REFERENCE

C6 H13 N O5 . H2 O4 S MF

> CM1

CRN 7664-93-9 CMF H2 O4 S

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

```
L72
     ANSWER 5 OF 10 REGISTRY COPYRIGHT 2003 ACS
     22161-81-5 REGISTRY
CN
     Benzeneacetic acid, 3-benzoyl-.alpha.-methyl-, (.alpha.S)- (9CI)
      (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Benzeneacetic acid, 3-benzoyl-.alpha.-methyl-, (S)-
CN
     Hydratropic acid, m-benzoyl-, (+)- (8CI)
OTHER NAMES:
CN
      (+)-(S)-m-Benzoylhydratropic acid
CN
      (+)-2-(3-Benzoylphenyl)propionic acid
      (+)-3-Benzoyl-.alpha.-methylbenzeneacetic acid
CN
      (+)-3-Benzoylhydratropic acid
CN
CN
      (+)-Ketoprofen
CN
      (2S)-2-(3-Benzoylphenyl)propionic acid
      (S)-(+)-2-(3-Benzoylphenyl) propionic acid
CN
      (S)-2-(3-Benzoylphenyl)propionic acid
CN
CN
     (S) -Ketoprofen
CN
     Dexketoprofen
CN
     S(+)-Ketoprofen
FS
     STEREOSEARCH
     C16 H14 O3
MF
CI
     COM
LC
                   ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*,
       BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, CEN, CHEMCATS,
       CHEMINFORMRX, CIN, CSCHEM, DDFU, DRUGNL, DRUGPAT, DRUGU, DRUGUPDATES, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, PHAR, PROMT, RTECS*, TOXCENTER,
       USAN, USPATFULL
          (*File contains numerically searchable property data)
     Other Sources:
                        WHO
```

Absolute stereochemistry. Rotation (+).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

382 REFERENCES IN FILE CA (1962 TO DATE)

4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

383 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:19408

REFERENCE 2: 138:11225

REFERENCE 3: 137:379776

REFERENCE 4: 137:375140

REFERENCE 5: 137:261988

REFERENCE 6: 137:257005

REFERENCE 7: 137:241443

REFERENCE 8: 137:162942

REFERENCE 9: 137:124232

REFERENCE 10: 137:108341

L72 ANSWER 6 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 14999-43-0 REGISTRY

CN D-Glucose, 2-amino-2-deoxy-, sulfate (2:1) (salt) (8CI, 9CI) (CA INDEX NAME)

OTHER NAMES:

CN 2-Amino-2-deoxy-.beta.-D-glucose sulfate

CN Tiocondramina

FS STEREOSEARCH

MF C6 H13 N O5 . 1/2 H2 O4 S

CI COM

LC STN Files: CA, CAPLUS, CHEMCATS, CHEMLIST, CSCHEM Other Sources: EINECS**

(**Enter CHEMLIST File for up-to-date regulatory information)

CM 1

CRN 7664-93-9 CMF H2 O4 S

CM 2

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

3 REFERENCES IN FILE CA (1962 TO DATE)

3 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 127:166636

REFERENCE 2: 107:218005

REFERENCE 3: 66:85991

L72 ANSWER 7 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 14257-69-3 REGISTRY

CN .beta.-D-Glucopyranose, 2-amino-2-deoxy- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Glucopyranose, 2-amino-2-deoxy-, .beta.-D- (8CI)

OTHER NAMES:

CN .beta.-D-Glucosamine

CN 2-Amino-2-deoxy-.beta.-D-glucopyranose

FS STEREOSEARCH

DR 28905-10-4

MF C6 H13 N O5

CI COM

LC STN Files: AGRICOLA, BEILSTEIN*, BIOBUSINESS, BIOSIS, CA, CAPLUS, CASREACT, CHEMINFORMRX, HODOC*, MRCK*, SPECINFO, TOXCENTER, USPATFULL (*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

50 REFERENCES IN FILE CA (1962 TO DATE)

10 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

50 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 137:201480

REFERENCE 2: 137:168260

REFERENCE 3: 136:216348

REFERENCE 4: 135:235349

REFERENCE 5: 135:149658

REFERENCE 6: 134:39012

REFERENCE 7: 133:59007

REFERENCE 8: 133:5384

REFERENCE 9: 131:161649

REFERENCE 10: 131:130186

L72 ANSWER 8 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 14131-63-6 REGISTRY

CN .beta.-D-Glucopyranose, 2-amino-2-deoxy-, hydrochloride (9CI) (CA INDEX NAME)

OTHER NAMES:

CN .beta.-Glucosamine hydrochloride

CN 2-Amino-2-deoxy-.beta.-D-glucopyranose hydrochloride

FS STEREOSEARCH

DR 140400-23-3

MF C6 H13 N O5 . C1 H

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, GMELIN*, HODOC*, SPECINFO, TOXCENTER

(*File contains numerically searchable property data)

CRN (14257-69-3)

Absolute stereochemistry.

● HCl

19 REFERENCES IN FILE CA (1962 TO DATE)

19 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 131:254291

REFERENCE 2: 131:228899

REFERENCE 3: 131:152961

REFERENCE 4: 129:224901

REFERENCE 5: 120:190966

REFERENCE 6: 119:96006

REFERENCE 7: 116:194737

REFERENCE 8: 107:237258

REFERENCE 9: 100:44337

REFERENCE 10: 99:158803

L72 ANSWER 9 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 14131-62-5 REGISTRY

CN .alpha.-D-Glucopyranose, 2-amino-2-deoxy-, hydrochloride (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Glucopyranose, 2-amino-2-deoxy-, hydrochloride, .alpha.-D- (8CI)

OTHER NAMES:

CN .alpha.-Glucosamine hydrochloride

CN 2-Amino-2-deoxy-.alpha.-D-glucopyranose hydrochloride

FS STEREOSEARCH

MF C6 H13 N O5 . C1 H

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, CHEMINFORMRX, CSCHEM, GMELIN*, SPECINFO, TOXCENTER

(*File contains numerically searchable property data)

(6490 - 70 - 6)CRN

Absolute stereochemistry.

● HCl

33 REFERENCES IN FILE CA (1962 TO DATE)

33 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 137:232735

REFERENCE 2: 137:79182

REFERENCE 3: 136:159026

REFERENCE 4: 134:326004

REFERENCE 5: 130:296924

REFERENCE 6: 130:81741

REFERENCE 7: 125:276366 REFERENCE 8: 121:256173

REFERENCE 9: 119:96006

REFERENCE 10: 118:115974

L72 ANSWER 10 OF 10 REGISTRY COPYRIGHT 2003 ACS

RN 6490-70-6 REGISTRY

CN .alpha.-D-Glucopyranose, 2-amino-2-deoxy- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Glucopyranose, 2-amino-2-deoxy-, .alpha.-D- (8CI)

OTHER NAMES:

CN .alpha.-D-Glucosamine

FS STEREOSEARCH

DR 66141-43-3, 28905-11-5

MF C6 H13 N O5

CI COM

LC STN Files: BEILSTEIN*, BIOSIS, CA, CAPLUS, CASREACT, CHEMINFORMRX, GMELIN*, MRCK*, SPECINFO, TOXCENTER, USPATFULL

(*File contains numerically searchable property data)

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

45 REFERENCES IN FILE CA (1962 TO DATE)

3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

45 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 135:189215

REFERENCE 2: 134:340642

REFERENCE 3: 133:252635

REFERENCE 4: 132:206486

REFERENCE 5: 131:199916

REFERENCE 6: 130:81741

REFERENCE 7: 129:276168

REFERENCE 8: 128:321844

REFERENCE 9: 128:57507

REFERENCE 10: 127:95502

FILE 'EMBASE' ENTERED AT 11:20:59 ON 27 JAN 2003 COPYRIGHT (C) 2003 Elsevier Science B.V. All rights reserved.

FILE COVERS 1974 TO 16 Jan 2003 (20030116/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all tot 191

L91 ANSWER 1 OF 5 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.

AN **2001268073** EMBASE

TI Sulfate could mediate the therapeutic effect of **glucosamine** sulfate.

AU Hoffer L.J.; Kaplan L.N.; Hamadeh M.J.; Grigoriu A.C.; Baron M.

CS Dr. L.J. Hoffer, Lady Davis Inst. for Medica Res, Jewish General Hospital, 3755 Cote-Ste-Catherine Rd, Montreal, Que. H3T 1E2, Canada

SO Metabolism: Clinical and Experimental, (2001) 50/7 (767-770). Refs: 31

ISSN: 0026-0495 CODEN: METAAJ

CY United States

DT Journal; Article

FS 037 Drug Literature Index
031 Arthritis and Rheumatism
030 Pharmacology

LA English

SL English

AΒ Glucosamine sulfate is a controversial osteoarthritis remedy that is presumed to stimulate articular cartilage glycosaminoglycan synthesis by increasing glucosamine concentrations in the joint space. However, this is not plausible because even large oral doses of the product have no effect on serum glucosamine consertions. We propose instead that sulfate could mediate the clinical benefit attributed to this treatment. Sulfate is required for glycosaminoglycan synthesis, and unlike glucosamine, its serum level can be modified by dietary and other factors. In this study, we tested whether oral glucosamine sulfate increases serum sulfate concentrations and whether the sulfate concentration in the synovial fluid reflects that in the serum. The serum sulfate concentration of 7 normal subjects was 331 .+-. 21 .mu.mol/L before ingestion of 1.0 g ${\tt glucosamine}$ sulfate and 375 .+-. 17 .mu.mol/L 3 hours after (P < .05). Serum sulfate concentrations decreased from 325 .+-. 19 to 290 .+-. 19 .mu.mol/L when the same dose of glucosamine sulfate was ingested with 1.0 g of the analgesic drug acetaminophen, which is largely metabolized by sulfation (P < .05). Unlike **glucosamine** sulfate, oral sodium sulfate did not significantly increase the serum sulfate concentration. Synovial fluid and serum sulfate concentrations were closely similar when measured in 15 patients undergoing diagnostic needle aspiration of a knee effusion (r = .99, slope = .97, P < .0001). These results do not prove that glucosamine sulfate improves osteoarthritis, but considered with other data, they do provide a plausible biochemical mechanism for its reported beneficial effects. This hypothesis is clinically relevant because it predicts that nonsulfate salts of glucosamine will be ineffective and that renal function, diet, and concurrent acetaminophen therapy could confound clinical trials of this therapy. Copyright .COPYRGT. 2001 by W.B. Saunders Company.

CT Medical Descriptors:
human
controlled study
clinical article
male

```
female
 drug effect
 osteoarthritis: DT, drug therapy
 articular cartilage
 glycosaminoglycan metabolism
 drug potentiation
 drug synovial fluid level
 drug blood level
 dietary intake
 concentration (parameters)
 dose response
 sulfation
drug metabolism
 joint effusion: DI, diagnosis
joint effusion: DT, drug therapy
needle biopsy
prediction
kidney function
article
priority journal
Drug Descriptors:
  *glucosamine sulfate: DT, drug therapy
  *glucosamine sulfate: PD, pharmacology
  *glucosamine sulfate: CR, drug concentration
  *glucosamine sulfate: IT, drug interaction
  *glucosamine sulfate: CB, drug combination
  *glucosamine sulfate: PO, oral drug administration
  *glucosamine sulfate: DO, drug dose
  *glucosamine sulfate: CM, drug comparison
  *glucosamine sulfate: PK, pharmacokinetics
*sulfate
glycosaminoglycan: EC, endogenous compound
  paracetamol: CB, drug combination
  paracetamol: PK, pharmacokinetics
  paracetamol: DT, drug therapy
sodium sulfate: PO, oral drug administration
sodium sulfate: CM, drug comparison
sodium sulfate: PD, pharmacology
(glucosamine sulfate) 29031-19-4; (sulfate)
14808-79-8; (paracetamol) 103-90-2; (sodium sulfate) 7757-82-6
SISU Enterprises (Canada)
ANSWER 2 OF 5 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.
2001252379 EMBASE
Current concepts regarding pharmacologic treatment of rheumatoid and
osteoarthritis.
Wildy K.S.; Wasko M.C.M.
Dr. M.C.M. Wasko, Division of Rheumatology, 3500 Terrace Street, BST South
700, Pittsburgh, PA 15261, United States
Hand Clinics, (2001) 17/2 (321-338).
Refs: 76
ISSN: 0749-0712 CODEN: HACLEO
United States
Journal; General Review
031
        Arthritis and Rheumatism
037
        Drug Literature Index
038
        Adverse Reactions Titles
English
English
Treating patients with osteoarthritis (OA) and rheumatoid arthritis (RA)
remains challenging; however, new agents offer the chance for an improved
quality of life. As an alternative to traditional nonsteroidal
anti-inflammatories, cyclooxygenase-2 inhibitors provide pain relief for
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LA

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AB

OA and RA patients with possible fewer side effects. Otherwise, OA patients may opt for topical agents, injections, or supplements. Rheumatoid arthritis research has led to an improved understanding of the inflammatory cascade and an appreciation of the early tissue destruction. A new treatment philosophy has thus emerged along with the development of new biologic agents; the latter, along with combination therapy and a new disease modifying antirheumatic drug, leflunomide, have greatly expanded the chances for disease control in RA patients. Medical Descriptors: *rheumatoid arthritis: DT, drug therapy *rheumatoid arthritis: ET, etiology *osteoarthritis: DT, drug therapy *osteoarthritis: ET, etiology drug effect enzyme activity enzyme blood level drug mechanism treatment outcome dose response risk factor protein expression disease severity drug indication dyspepsia: SI, side effect abdominal pain: SI, side effect analgesia risk assessment nausea and vomiting: SI, side effect headache: SI, side effect bone marrow suppression: SI, side effect drug potency rash: SI, side effect alopecia: SI, side effect human clinical trial review Drug Descriptors: *nonsteroid antiinflammatory agent: AE, adverse drug reaction *nonsteroid antiinflammatory agent: DT, drug therapy *nonsteroid antiinflammatory agent: TO, drug toxicity *nonsteroid antiinflammatory agent: PD, pharmacology *corticosteroid: AE, adverse drug reaction *corticosteroid: DT, drug therapy *corticosteroid: PD, pharmacology *corticosteroid: AR, intraarticular drug administration *corticosteroid: PO, oral drug administration *cyclooxygenase 2 inhibitor: ĀE, adverse drug reaction *cyclooxygenase 2 inhibitor: DT, drug therapy *cyclooxygenase 2 inhibitor: PD, pharmacology *antirheumatic agent: AE, adverse drug reaction *antirheumatic agent: CT, clinical trial *antirheumatic agent: DT, drug therapy *antirheumatic agent: PD, pharmacology cyclooxygenase 1: EC, endogenous compound cyclooxygenase 2: EC, endogenous compound arachidonic acid: EC, endogenous compound leukotriene: EC, endogenous compound phospholipase A2: EC, endogenous compound thromboxane A2: EC, endogenous compound ketoprofen: DT, drug therapy flurbiprofen: DT, drug therapy indometacin: DT, drug therapy

piroxicam: DT, drug therapy

CT

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naproxen: DT, drug therapy
   ibuprofen: DT, drug therapy
   diclofenac: DT, drug therapy
   etodolac: DT, drug therapy
 meloxicam: DT, drug therapy
rofecoxib: AE, adverse drug reaction
 rofecoxib: DT, drug therapy
celecoxib: AE, adverse drug reaction
celecoxib: DT, drug therapy
hyaluronic acid: DT, drug therapy
   glucosamine sulfate: DT, drug therapy
hydroxychloroquine: AE, adverse drug reaction
hydroxychloroquine: CT, clinical trial
   hydroxychloroquine: CB, drug combination
hydroxychloroquine: DT, drug therapy
hydroxychloroquine: PD, pharmacology
salazosulfapyridine: AE, adverse drug reaction
salazosulfapyridine: CT, clinical trial
   salazosulfapyridine: CB, drug combination
salazosulfapyridine: DT, drug therapy
salazosulfapyridine: PD, pharmacology
methotrexate: AE, adverse drug reaction
methotrexate: CT, clinical trial
  methotrexate: CB, drug combination
methotrexate: DT, drug therapy
methotrexate: PD, pharmacology
methotrexate: PO, oral drug administration
cyclosporin: AE, adverse drug reaction
  cyclosporin: CB, drug combination
cyclosporin: DT, drug therapy
cyclosporin: PD, pharmacology
leflunomide: AE, adverse drug reaction
leflunomide: DT, drug therapy
leflunomide: PD, pharmacology
etanercept: CT, clinical trial
etanercept: DT, drug therapy
etanercept: PD, pharmacology
infliximab: CT, clinical trial
infliximab: DT, drug therapy
infliximab: PD, pharmacology
glucocorticoid: AE, adverse drug reaction
glucocorticoid: DT, drug therapy
  capsaicin: AE, adverse drug reaction
  capsaicin: DT, drug therapy
  capsaicin: TP, topical drug administration
azathioprine: DT, drug therapy
  actron
  diclofenac potassium
(arachidonic acid) 506-32-1, 6610-25-9, 7771-44-0; (phospholipase A2) 9001-84-7; (thromboxane A2) 57576-52-0; (ketoprofen)
22071-15-4, 57495-14-4; (flurbiprofen) 5104-49-4; (indometacin)
53-86-1, 74252-25-8, 7681-54-1; (piroxicam) 36322-90-4; (naproxen)
22204-53-1, 26159-34-2; (ibuprofen) 15687-27-1; (diclofenac) 15307-79-6, 15307-86-5; (etodolac) 41340-25-4; (meloxicam) 71125-38-7; (rofecoxib) 162011-90-7, 186912-82-3; (celecoxib) 169590-42-5;
(hyaluronic acid) 31799-91-4, 9004-61-9, 9067-32-7; (glucosamine sulfate) 29031-19-4; (hydroxychloroquine) 118-42-3, 525-31-5;
(salazosulfapyridine) 599-79-1; (methotrexate) 15475-56-6, 59-05-2,
7413-34-5; (cyclosporin) 79217-60-0; (leflunomide) 75706-12-6;
(etanercept) 185243-69-0, 200013-86-1; (infliximab) 170277-31-3;
(capsaicin) 404-86-4; (azathioprine) 446-86-6; (diclofenac potassium)
15307-81-0
(1) Remicade; Orudis; Actron; Ansaid; Indocin; Feldene; Naprosyn; Motrin;
```

RN

CN

```
Advil; Voltaren; Cataflam; Lodine; Mobic; Vioxx; Celebrex; Plaquenil;
     Azulfidine; Arava
CO
     (1) Centecor (United States)
L91 ANSWER 3 OF 5 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.
     2000063108 EMBASE
ΑN
TI
     Glucosamine in the treatment of osteoarthritis.
ΑU
     Delafuente J.C.
     J.C. Delafuente, 410 North 12th Street, Richmond, VA 23298-0533, United
CS
     States
SO
     Rheumatic Disease Clinics of North America, (2000) 26/1 (1-11).
     Refs: 21
     ISSN: 0889-857X CODEN: RDCAEK
     United States
CY
DΤ
     Journal; General Review
     031
             Arthritis and Rheumatism
     033
             Orthopedic Surgery
     037
             Drug Literature Index
     038
             Adverse Reactions Titles
LA
     English
ST
     English
AΒ
     Glucosamine sulfate, a constituent of cartilage, is evaluated
     for the treatment of osteoarthritis. Available data suggest that
     glucosamine decreases pain and improves function in
     osteoarthritis. Most of the glucosamine studies have
     methodological flaws or used parenteral formulations, making
     their data difficult to extrapolate into clinical practice.
     Glucosamine sulfate is shown to be as good as ibuprofen
     for osteoarthritis of the knee. Better designed clinical trials of
     glucosamine are needed to identify its role in the pharmacotherapy
     of osteoarthritis.
CT
    Medical Descriptors:
     *knee osteoarthritis
     *pain assessment
    functional disease
    drug efficacy
    drug mechanism
    articular cartilage
    drug bioavailability
    dose response
    heartburn: SI, side effect
    epigastric pain: SI, side effect
    nausea: SI, side effect
    diet supplementation
    human
    controlled study
    review
    priority journal
    Drug Descriptors:
      *glucosamine sulfate: AE, adverse drug reaction
      *glucosamine sulfate: AD, drug administration
      *glucosamine sulfate: CM, drug comparison
      *glucosamine sulfate: DO, drug dose
      *glucosamine sulfate: PK, pharmacokinetics
      *glucosamine sulfate: PD, pharmacology
      *glucosamine sulfate: AR, intraarticular drug administration
      *glucosamine sulfate: IM, intramuscular drug administration
      *glucosamine sulfate: IV, intravenous drug administration
      *glucosamine sulfate: PO, oral drug administration
      ibuprofen: CM, drug comparison
      ibuprofen: DO, drug dose
      ibuprofen: PD, pharmacology
    placebo
```

chlorbutol: CM, drug comparison chlorbutol: IM, intramuscular drug administration piperazine: CM, drug comparison piperazine: IM, intramuscular drug administration chondroitin sulfate RN (glucosamine sulfate) 29031-19-4; (ibuprofen) 15687-27-1; (chlorbutol) 57-15-8; (piperazine) 110-85-0, 142-63-2, 142-64-3, 16832-43-2, 6094-40-2; (chondroitin sulfate) 9007-28-7, 9082-07-9 L91 ANSWER 4 OF 5 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V. AN **94150643** EMBASE DN 1994150643 TΤ Glucosamine sulfate compared to ibuprofen in osteoarthritis of the knee. ΑU Muller-Fassbender H.; Bach G.L.; Haase W.; Rovati L.C.; Setnikar I. CS Department of Clinical Pharmacology, Rotta Research Laboratorium, Via Valosa di Sopra 7,20052 Monza, Italy Osteoarthritis and Cartilage, (1994) 2/1 (61-69). SO ISSN: 1063-4584 CODEN: OSCAEO CY United Kingdom DTJournal; Article FS 019 Rehabilitation and Physical Medicine 031 Arthritis and Rheumatism 030 Pharmacology 037 Drug Literature Index 038 Adverse Reactions Titles LΑ English SL English AΒ Glucosamine sulfate is able to stimulate proteoglycan synthesis by chondrocytes and has mild anti-inflammatory properties. In clinical trials, glucosamine sulfate was more effective than placebo in controlling the symptoms of osteoarthritis (OA). In order to better characterize this therapeutic activity, we conducted a randomized, double-blind, parallel-group study of glucosamine sulfate 500 mg t.i.d. vs ibuprofen 400 mg t.i.d., orally for 4 weeks. The study included 200 hospitalized patients with active OA of the knee, symptoms for at least 3 months and a Lequesne's index of at least 7 points. Patients were evaluated weekly. Response was defined as a reduction in the Lequesne's index by at least 2 points if the enrolment value was higher than 12 points, or by at least 1 point if the enrolment value was 12 or less points, together with a positive overall assessment by the investigator. The improvement tended to be sooner under ibuprofen (48% responders vs 28% after the 1st treatment week; P = 0.06, Fisher's Exact test), but there was no difference from the 2nd week onward, with a success rate of 52% in the ibuprofen group and of 48% in the glucosamine group (P = 0.67) at the end of treatment. The average Lequesne's index at enrolment was around 16 points and decreased by over 6 points in both groups, again with the above described trend. On the other hand, 35% of patients on ibuprofen reported adverse events, mainly of gastrointestinal origin, vs 6% adverse events with glucosamine (P < 0.001, Fisher's Exact test). The number of adverse event related drop-outs was different between the two groups (7% vs 1%, respectively; P = 0.035). Glucosamine sulfate was therefore as effective as ibuprofen on symptoms of knee OA. These data confirm glucosamine sulfate as a safe symptomatic Slow Acting Drug for OA. CTMedical Descriptors: *knee osteoarthritis: DT, drug therapy *knee osteoarthritis: TH, therapy adult aged article

clinical trial controlled study double blind procedure fatigue: SI, side effect female flushing gastrointestinal disease: SI, side effect major clinical study male oral drug administration priority journal pruritus: SI, side effect randomized controlled trial rash: SI, side effect side effect Drug Descriptors: *glucosamine sulfate: DT, drug therapy *glucosamine sulfate: AE, adverse drug reaction *glucosamine sulfate: CM, drug comparison *glucosamine sulfate: CT, clinical trial *ibuprofen: DT, drug therapy *ibuprofen: AE, adverse drug reaction *ibuprofen: CM, drug comparison *ibuprofen: CT, clinical trial RN (glucosamine sulfate) 29031-19-4; (ibuprofen) 15687-27-1 ANSWER 5 OF 5 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V. **80119580** EMBASE DN 1980119580 Biochemical gastroprotection from acute ulceration induced by aspirin and TΤ related drugs. ΑU Rainsford K.D.; Whitehouse M.W. CS Biochem. Dept., Univ. Tasmania Med. Sch., Hobart, Australia Biochemical Pharmacology, (1980) 29/9 (1281-1289). SO CODEN: BCPCA6 CY United Kingdom DΤ Journal FS 037 Drug Literature Index 030 Pharmacology LA English Adjuncts that serve as: (a) buffers to neutralize drug acidity, and/or (b) AB solubilizers of acidic drugs, or (c) certain nutrients (e.g. glucose, acetate), considerably reduced the gastric mucosal injury induced by orally administered aspirin (and other non-steroidal anti-inflammatory drugs) in stressed and starved rats. Gastroprotection against aspirin and related drugs was obtained by supplying the mucosa with glucose with intermediates or precursors of the tricarboxylic acid cycle (that may be absorbed directly from the gastric lumen). Glucose alone was not sufficiently gastroprotective. Gastroprotection with tricarboxylic acid

related drugs was obtained by supplying the mucosa with glucose with intermediates or precursors of the tricarboxylic acid cycle (that may be absorbed directly from the gastric lumen). Glucose alone was not sufficiently gastroprotective. Gastroprotection with tricarboxylic acid cycle precursors given with glucose appears to be due to the effects of these nutrients in restoring ATP synthesis in the gastric mucosa. D-glutamate and D-aspartate were deaminated by homogenates prepared from saline-washed rat fundic mucosa (yielding .alpha.-oxo acids for the tricarboxylic acid cycle). These amino acids could be substituted for the L-forms in combination with glucose to yield gastroprotection from damage by aspirin. Studies in domestic pigs (a species with a pseudo-human stomach) established that acute and chronic oral administration of the aspirin+acetate+glucose combination (1:3:3 molar proportions) was less irritating to the gastric mucosa than aspirin alone. These results were confirmed in acute studies in monkeys. Sodium and potassium salts were superior to calcium and ammonium salts as the

buffer component in these improved (i.e. less gastrotoxic) aspirin formulations tested in rats. Bicarbonate was not effective in preventing aspirin gastrotoxicity in stressed-sensitized rats, but is effective in non-stressed rats. CTMedical Descriptors: *3,5 dibromoacetylsalicylic acid *cell metabolism *formate sodium *malate sodium *maleate sodium *malonate sodium *stomach mucosa *stomach ulcer adverse drug reaction pathogenesis ph stress etiology preliminary communication animal experiment oral drug administration stomach swine rat drug comparison Drug Descriptors: *3 hydroxybutyric acid *acetic acid *acetylsalicylic acid *alanine *ammonium acetate *antiinflammatory agent *arginine *aspartic acid *benzoic acid *butyric acid *calcium acetate *citrate trisodium *citric acid *disodium hydrogen phosphate *fructose *galactose *glucose *glutamine *glycerol *glycine *lactate sodium *lactose *lysine *n acetylglucosamine *sodium dihydrogen phosphate *pyruvate sodium *ribose *sucrose RN (3 hydroxybutyric acid) 300-85-6; (acetic acid) 127-08-2, 127-09-3, 64-19-7, 71-50-1; (acetylsalicylic acid) 493-53-8, 50-78-2, 53663-74-4, 53664-49-6, 63781-77-1; (alanine) 56-41-7, 6898-94-8; (ammonium acetate) 631-61-8; (arginine) 1119-34-2, 15595-35-4, 7004-12-8, 74-79-3; (aspartic acid) 56-84-8, 6899-03-2; (benzoic acid) 532-32-1, 582-25-2, 65-85-0, 766-76-7; (butyric acid) 107-92-6, 156-54-7, 461-55-2; (calcium acetate) 62-54-4; (citrate trisodium) 6132-04-3, 68-04-2; (citric acid) 126-44-3, 5949-29-1, 77-92-9, 8002-14-0; (disodium hydrogen phosphate) 7558-79-4; (fructose) 30237-26-4, 57-48-7, 7660-25-5, 77907-44-9; (galactose)

26566-61-0, 50855-33-9, 59-23-4; (glucose) 50-99-7, 84778-64-3; (glutamine) 56-85-9, 6899-04-3; (glycerol) 56-81-5; (glycine) 56-40-6, 6000-43-7, 6000-44-8; (lactate sodium) 72-17-3; (lactose) 10039-26-6, 16984-38-6, 63-42-3, 64044-51-5; (lysine) 56-87-1, 6899-06-5, 70-54-2; (nacetylglucosamine) 7512-17-6; (sodium dihydrogen phosphate) 7558-80-7, 7632-05-5; (pyruvate sodium) 113-24-6; (ribose) 34466-20-1, 50-69-1, 93781-19-2; (sucrose) 122880-25-5, 57-50-1

=> fil wpix FILE 'WPIX' ENTERED AT 11:43:53 ON 27 JAN 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

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L128 ANSWER 1 OF 12 WPIX (C) 2003 THOMSON DERWENT

AN 2002-616446 [66] WPIX

DNN N2002-487716 DNC C2002-174257

TI Method, useful for treating musculoskeletal disorders, comprises topical application of a composition comprising penetration enhancers and an antiinflammatory agent in a gel vehicle.

DC A96 B04 B05 P34

IN PETRUS, E J

PA (ADME-N) ADVANCED MEDICAL INSTR

CYC 1

PI US 6399093 B1 20020604 (200266)* 9p A61L015-16

ADT US 6399093 B1 US 1999-314829 19990519

PRAI US 1999-314829 19990519

IC ICM A61L015-16

AB US 6399093 B UPAB: 20021014 NOVELTY - Method (I), comprises topical application of a composition (II) comprising:

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fonda - 09 / 964178
            (A) penetration enhancers from alcohols, polyols, sulfoxides, esters,
      ketones, amides, oleates, surfactants, alkanoic acids, lactam compounds,
      alkanols or dialkylamino acetates; and
           (B) an antiinflammatory agent from non-steroidal antiinflammatory
      agent or colchicine (0.1 - 25\% \text{ of } (I)), in a gel vehicle.
           DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for a
      composition (II) for use in the method.
           ACTIVITY - Osteopathic; Muscular; Antiinflammatory; Analgesic.
           No biological data available.
           MECHANISM OF ACTION - None given.
           USE - (I) is useful for treating musculoskeletal disorders (claimed)
      and providing antiinflammatory and analgesic benefits.
      Dwg.0/0
      CPI GMPI
      AB; DCN
     CPI: A99-A; B04-A10; B06-A01; B06-A03; B06-E05; B08-D01; B10-A10;
           B10-B02A; B10-C04C; B14-C01; B14-C03
TECH
                     UPTX: 20021014
     TECHNOLOGY FOCUS - BIOLOGY - Preferred Composition: (II) may further
     include analgesics, antioxidants, anti-infective agents, adjuvants,
     anthocyanidins, proanthocyanidins, muscle relaxants, nitric oxide synthase
     inhibitors, methyl-sulfonyl-methane, S-adenosyl-methionine, zinc
     compounds, aloe vera extract, amino sugars, glycosaminoglycans, manganese,
     magnesium, boron or herbal derivatives (claimed).
      (II) comprises 0.2 - 0.5 mg of colchicine and 2 - 6% of etodolac,
     ibuprofen or diclofenac.
ABEX
     EXAMPLE - A typical joint analgesic gel comprises ibuprofen
     (5%), carboxyvinyl polymers (2%), aloe vera gel (1%), propylene glycol
     (20%), glucosamine sulfate (20%), methyl-sulfonyl-methane (10%),
     ethanol (10%), triethanolamine (1%), zinc sulfate (1%), methyl paraben
     (0.1\%), propyl paraben (0.02\%) and water (29.9\%).
L128 ANSWER 2 OF 12 WPIX (C) 2003 THOMSON DERWENT
     2002-372086 [40]
                        WPIX
     C2002-105331
     Composition used for treating pain comprises glucosamine
     material and analgesic compound.
     COWAN, A; RAFFA, R; TALLARIDA, R
(UTEM) UNIV TEMPLE
CYC 97
     WO 2002026239 A1 20020404 (200240)* EN
                                               23p
                                                      A61K031-70
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TR TZ UG ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR
            KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PH PL PT RO
            RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW
     US 2002058642 A1 20020516 (200240)
                                                     A61K031-7008
     AU 2001092929 A 20020408 (200252)
                                                      A61K031-70
    WO 2002026239 A1 WO 2001-US29606 20010921; US 2002058642 A1 Provisional US
     2000-235405P 20000926, US 2001-964178 20010925; AU 2001092929 A AU
     2001-92929 20010921
FDT AU 2001092929 A Based on WO 200226239
PRAI US 2000-235405P 20000926; US 2001-964178
                                                 20010925
     ICM A61K031-70; A61K031-7008
     ICS A61K031-192
    WO 200226239 A UPAB: 20020626
    NOVELTY - Dosage form (I) comprises a glucosamine material (II)
    and an analgesic compound (III) in a weight ratio that results in an
    analgesic effect at least as great as that expected for the analgesic
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FS

FΑ

MC

DNC

TI

DC IN

PΑ

PΤ

AB

alone at that dosage.

fonda - 09 / 964178 ACTIVITY - Analgesic. MECHANISM OF ACTION - None given in the source material. USE - Useful for the control of pain caused by a wide variety of disorders e.g. colds and influenza, arthritis, headache, toothache, dysmenorrhea, surgery and muscle and joint pain. ADVANTAGE - The components of (I) give a synergistic analgesic DESCRIPTION OF DRAWING(S) - The drawing shows the analgesic effect of ibuprofen alone and a combination of ibuprofen and glucosamine sulfate on acetylcholine-induced abdominal constrictions in mice. Dwg.1/3FS CPI FΑ AB; GI; DCN MC CPI: B10-A07; B10-C04B; B10-C04C; B14-C01; B14-C09; B14-S09 TECH UPTX: 20020626 TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Materials: (III) Comprises a non steroidal antiinflammatory drug, preferably a propionic acid analgesic, ibuprofen or ketoprofen. (II) Is glucosamine or its salts, alpha- or beta-glucosamine, N-acetyl glucosamine, glucosamine sulfate or glucosamine hydrochloride. Preferred Composition: The ratio of (II) to (III) is at least 1:2, especially 1:2-10:1. (I) Also contains anti arthritic, antihistamine, muscle relaxant, sleep aid, decongestant and/or bronchodilator. ABEX ADMINISTRATION - The dosage is 10-6000 mg/kg/day orally, parenterally or topically. EXAMPLE - Glucosamine sulfate (250 mg) and ibuprofen (27.8 mg) were combined with water (10 ml) containing 2 drops of Tween (RTM; dispersant) to give a composition with a glucosamine to ibuprofen weight ratio of 5.2:1. In tests on mice, various combinations of glucosamine sulfate and analgesic were administered and the pain relief recorded using an acetylcholine-induced abdominal constriction model. The results showed that ibuprofen alone exhibited an ED50 value of 26 mg/kg, but the combination of ibuprofen with glucosamine sulfate (9:1 ratio) gave an $ED\bar{5}0$ value of 2.08 mg/kg. L128 ANSWER 3 OF 12 WPIX (C) 2003 THOMSON DERWENT **2002-179010** [23] WPIX 2002-082384 [01]; 2002-082386 [01]; 2002-082387 [01]; 2002-121526 [11]; 2002-381872 [36] DNC C2002-055485 Composition useful for treatment of pain comprises a pharmaceutical, nutraceutical and a base. GELBER, D; KLEINBERGER, R (GELB-I) GELBER D; (KLEI-I) KLEINBERGER R CYC US 2002006445 A1 20020117 (200223)* 16p A61K035-78 ADT US 2002006445 Al Provisional US 2000-184351P 20000223, US 2001-754204 20010105 PRAI US 2000-184351P 20000223; US 2001-754204 20010105 ICM A61K035-78

AB US2002006445 A UPAB: 20020701

AN

CR

TΤ

DC ΙN

PΑ

PΙ

IC

NOVELTY - A composition comprises at least one pharmaceutical (P1), at least one nutraceutical (N1) for treating pain resulting from an inflammatory response and a base.

ACTIVITY - Analgesic; virucide, antiallergic; antimigraine; gynecological; antiinflammatory; antiarthritic; antipyretic; antitussive. MECHANISM OF ACTION - None given.

USE - For the treatment of a patient suffering from pain resulting from an inflammatory response (claimed); also for the treatment of predetermined symptoms of an ailment; for the treatment of symptoms of colds, flu, allergies or sinus discomfort; for discomfort associated with heartburn, general body aches, headaches, migraines, menstruation, joint discomfort and arthritis. Also useful for the treatment of immune response (e.g. sinus congestion; red, itchy or watery eyes; and sneezing) resulting from exposure to atmospheric pollutants or allergens. Useful in healing process as well as preventing future maladies.

ADVANTAGE - The composition comprising combinations of pharmaceutical and nutraceutical increases the beneficial effects of the pharmaceutical utilized. The composition not only treats a current ailment more effectively, but also functions to prevent the recurrence of illness. Dwg.0/0

FS CPI

ro CPI

FA AB; DCN

MC CPI: B03-F; B04-A07E; B04-A08; B04-A10; B04-C02; B05-A03A; B06-D01; B06-F03; B07-A01; B07-D02; B07-D08; B10-A10; B10-B02; B10-C04; B10-D03; B10-F02; B14-A02; B14-C01; B14-C03; B14-C04; B14-C09; B14-G02A; B14-K01B; B14-N14; B14-S08

TECH

UPTX: 20020411
TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Components: (P1) is selected from a group consisting of pain relieving and/or anti-inflammatory agents (preferably acetaminophen and/or non-steroidal anti-inflammatory drug (NSAID)). (N1) is selected from immune boosters, anti-oxidants, a liver protectant and/or joint relief agents. The nutraceutical immune booster is selected from zinc and its salt and at least one herb selected from Echinacea, Sambucus and/or Goldenseal. The nutraceutical anti-oxidant is selected from a bioflavonoid, at least one herbal extract containing at least one bioflavonoid, ascorbic acid and its salt, garlic and its extract, green tea and its extracts, and/or at least one herb selected from Astragalus. The nutraceutical liver protectant is milk thistle. The nutraceutical joint relief agent is selected from glucosamine, glucosamine sulfate, chondroitin, chondroitin sulfate and its salts.

ABEX

SPECIFIC COMPOUNDS - Acetaminophen, diclophenac, fenflofenac, aspirin, indomethacin, sulindac, tolmetin, ibuprofen, ketoprofen, fenoprofen, flurbiprofen, naproxen, meclofenamic acid, flufenamic acid, piroxicam, tenoxicam, meloxicam, celicoxib, roficoxib and/or nabumetone are specifically claimed as (P1).

 $\label{lem:administered} \mbox{\sc administered orally and topically onto nasal mucosa.}$

EXAMPLE - A solid composition comprised (mg) acetaminophen (60 - 1000, preferably 200 - 750 and more preferably 350 - 550), diphenhydramine (5 -100, preferably 10 - 50, more preferably 20 - 40), pseudoephedrine (5 - 100, preferably 10 - 75, more preferably 20 - 40), Echinacea purpurea (10 - 500, preferably 25 - 200, more preferably 50 - 100), Goldenseal (50 -200, preferably 75 - 150, more preferably 80 - 120), Elderberry (sambucol) (50-250, preferably 75-175, more preferably 100-150), garlic extract (50-200, preferably 75-150, more preferably 80-120), green teaextract (50 - 120, preferably 75 - 150, more preferably 80 - 120), astragalus (50 - 250, preferably 75 - 175, more preferably 100 - 150), zinc gluconate (0.1 - 15, preferably 0.5 - 10, more preferably 0.5 - 7.5 and ascorbic acid (50 - 1000, preferably 100 - 750, more preferably 200 -500). The composition is administered in the form of a capsule to a mammal, for treating the symptoms of a cold or flu every 4 - 6 hours to relieve pain and discomfort and to provide immune system stimulation. Echinacea purpurea and Astragalus assisted in boosting the immune system while the pharmaceutical components treat the symptoms associated with inflammatory and mucous accumulation.

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L128 ANSWER 4 OF 12 WPIX (C) 2003 THOMSON DERWENT
      2001-565929 [64]
                         WPIX
 CR
      2001-457762 [50]
 DNC
     C2001-167995
      Treatment of photophobia and phonophobia associated with migraine attack,
      comprises use of ibuprofen.
 DC
      A96 B05
 ΙN
     CODISPOTI, J R
PΑ
      (MCNI) MCNEIL-PPC INC
CYC
     31
PΙ
     CA 2326549
                    A1 20010524 (200164) * EN
                                                23p
                                                       A61K031-192
      CN 1298700
                    A 20010613 (200164)
                                                       A61K031-19
      EP 1129710
                   A2 20010905 (200164) EN
                                                       A61K031-192
          R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
             RO SE SI TR
     JP 2001192332 A 20010717 (200164)
KR 2001051939 A 20010625 (200172)
NZ 508292 A 20020830 (200265)
                                                 6p
                                                       A61K031-192
                                                       A61K031-192
                                                       A61K031-615
     CA 2326549 A1 CA 2000-2326549 20001123; CN 1298700 A CN 2000-128380
     20001123; EP 1129710 A2 EP 2000-310391 20001123; JP 2001192332 A JP
     2000-356516 20001122; KR 2001051939 A KR 2000-70461 20001124; NZ 508292 A
     NZ 2000-508292 20001120
    NZ 508292 A Div in NZ 518742
PRAI US 2000-709069
                      20001109; US 1999-449124
                                                   19991124
     ICM A61K031-19; A61K031-192; A61K031-615
          A61K031-535; A61P025-06
AB
          2326549 A UPAB: 20021010
     NOVELTY - A method for mitigation or treatment of photophobia and
     phonophobia associated with migraine comprises the use of
     ibuprofen (a), its salts (b) and/or its isomers (c).
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are included for:
           (i) a medicament for mitigating or treating photophobia and
     phonophobia associated with migraine comprising (a), (b) and/or (c);
           (ii) a composition for mitigating or treating photophobia and
     phonophobia associated with migraine comprising (a), (b) and/or (c); and
           (iii) a commercial package for mitigating or treating photophobia and
     phonophobia associated with migraine, which comprises (a), (b) and/or (c)
     together with instructions for use of the package.
          ACTIVITY - Antimigraine; analgesic; antiinflammatory.
          Of the 650 patients suffering from photophobia due to migraine some
     patients were given placebo, while others were given either 200 or 400 mg
     of ibuprofen. Periodic assessments of their photophobia were
     made by determining % of patients with severity reduced to zero. The
     results for ibuprofen (200 mg)/ibuprofen (400
     mg)/placebo-treated patients showed that % of patients with photophobia
     severity reduced to zero at a time (hours) of 2, 3 and 5 was 22/19/15;
     29/26/19 and 37/34/26 respectively.
          MECHANISM OF ACTION - None given.
          USE - For mitigating or treating photophobia and phonophobia
     associated with migraines (claimed).
          ADVANTAGE - The method utilizes the action of a single active
     ingredient, which is commercially available and does not cause undesired
     side effects.
     Dwg.0/2
FS
     CPI
FΑ
     AB; DCN
MC
     CPI: A12-V01; B10-C04C; B14-C01; B14-C03; B14-J01
TECH
                    UPTX: 20011105
     TECHNOLOGY FOCUS - INORGANIC CHEMISTRY - Preferred Component: (b) is an
     inorganic cation salt (preferably sodium, potassium, lithium, magnesium,
     calcium, cesium, ammonia, ferrous, zinc, manganous, aluminum, ferric or
     manganic).
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TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Components: (b) is an inorganic cation salt and/or an organic salt of (a) with primary, secondary, tertiary or quaternary amine (preferably triethylamine, tripropylamine, 2-dimethylaminoethanol, 2-diethylaminoethanol, lysine, arginine, histidine, caffeine, procain, N-ethyl piperidine, hydrabamine, choline, betaine, ethylenediamine, glucosamine, tris(hydroxymethyl)aminomethane, methylglycamine, theobromine, purine, piperazine, piperidine, or polyamine resin). The mixture of (a), (b) and (c) is preferably a mixture of (a) and its sodium salt. (c) is Ribuprofen and/or S-ibuprofen.

ABEX

ADMINISTRATION - The administration is oral. The amount of ibuprofen and/or its isomer is 100-800 (preferably 200-400) mg per dose. The salt of the ibuprofen is administered in a dosage of about 100-1700 mg per dose, whereas the mixture of ibuprofen, its isomer and its salt is administered in a dosage of about 100-1700 (preferably 200-1300) mg per dose.

EXAMPLE - None given.

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L128 ANSWER 5 OF 12 WPIX (C) 2003 THOMSON DERWENT
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ΑN **2001-457762** [50] WPIX

CR 2001-565929 [64]

DNC C2001-138476

Use of an ibuprofen for treating photophobia and phonophobia associated with a migraine attack.

DC B05

ΙN CODISPOTI, J R

PΑ (MCNI) MCNEIL-PPC INC; (JOHJ) JOHNSON & JOHNSON

CYC

PΙ AU 2000071733 A 20010531 (200150)* 13p A61K031-192 BR 2000005560 A 20010731 (200150) A61K031-192 KR 2001051939 A 20010625 (200172) A61K031-192 ZA 2000006885 A 20021030 (200282) 16p A61K000-00

AU 2000071733 A AU 2000-71733 20001121; BR 2000005560 A BR 2000-5560 ADT 20001124; KR 2001051939 A KR 2000-70461 20001124; ZA 2000006885 A ZA 2000-6885 20001123

PRAI US 1999-449124 19991124; US 2000-709069 20001109

ICM A61K000-00; A61K031-192

ICS A61P025-06

AU 200071733 A UPAB: 20021220 AΒ NOVELTY - Mitigation or treatment of photophobia and phonophobia associated with migraine involves the use of an ibuprofen (a), its salts (b) and/or its isomers (c).

ACTIVITY - Antimigraine; Analgesic; Antiinflammatory.

Of the 650 patients suffering from photophobia due to migraine some patients were given placebo, while others were given either 200 or 400~mgof ibuprofen. Periodic assessments of their photophobia were made by determining % of patients with severity reduced to zero. The results for ibuprofen (200 mg)/ibuprofen (400 mg)/placebo-treated patients showed that % of patients with photophobia severity reduced to zero at a time (hours) of 2,3 and 5 was 22/19/15; $29/26/\overline{19}$ and 37/34/26 respectively.

MECHANISM OF ACTION - None given.

USE - For mitigating or treating photophobia and phonophobia associated with migraines (claimed).

ADVANTAGE - The method utilizes the action of a single active ingredient, which is commercially available and does not cause undesired side effects.

Dwq.0/2

FS CPI

FA AB; DCN MC CPI: **B10-C04C**; **B14-C01**; B14-N02; B14-N03 TECH UPTX: 20010905

TECHNOLOGY FOCUS - INORGANIC CHEMISTRY - Preferred Component: (b) is an inorganic cation salt (preferably sodium, potassium, lithium, magnesium, calcium, cesium, ammonia, ferrous, zinc, manganous, aluminum, ferric or manganic).

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred Components: (b) is an inorganic cation salt; and/or an organic salt of (a) with primary, secondary, tertiary or quaternary amine (preferably triethylamine, tripropylamine, 2-dimethylaminoethanol, 2-diethylaminoethanol, lysine, arginine, histidine, caffeine, procain, N-ethyl piperidine, hydrabamine, choline, betaine, ethylenediamine, glucosamine, TRIS (hydroxymethyl)aminomethane, methylglycamine, theobromine, pruine, piperazine, and/or piperidine). The mixture of (a), (b) and (c) is preferably a mixture of (a) and its sodium salt. (c) is R-ibuprofen and/or S-ibuprofen.

TECHNOLOGY FOCUS - POLYMERS - The organic salt is a polyamine resin.

ABEX

ADMINISTRATION - The administration is oral. The amount of ibuprofen and/or its isomer is 100 - 800 (preferably 200 - 600, especially 200 - 400) mg per dose. The salt of the ibuprofen is administered in a dosage of about 100 - 1700 mg per dose, whereas the mixture of ibuprofen, its isomer and its salt is administered in a dosage of about 100 - 1700 (preferably 200 - 1300) mg per dose (all claimed).

EXAMPLE - None given.

L128 ANSWER 6 OF 12 WPIX (C) 2003 THOMSON DERWENT

AN 2001-182859 [18] WPIX

DNC C2001-054566

TI Treatment of pain due to degenerative joint diseases associated with inflammation in cats and dogs, using flupirtine as potent analgesic having low side-effect potential.

DC B05 C02 C03

IN ENDLER, G; LEHMANN, H; LOBISCH, M; SZELENYI, I; LOBICH, M

PA (ASTA) ASTA MEDICA AG; (FARB) BAYER AG; (DRED) AWD PHARMA GMBH & CO KG CYC 56

PI WO 2001008682 A2 20010208 (200118)* DE 20p A61K031-44 RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: AU BG BR BY CA CN CZ EE GE HR HU ID IL IN IS JP KG KR KZ LT LV MK MX NO NZ PL RO RU SG SI SK TR UA UZ YU ZA

CA 2314746 A1 20010203 (200119) EN A61K031-44
AU 2000072717 A 20010219 (200129) A61K031-44
NO 2002000364 A 20020123 (200231) A61K031-44
BR 2000012942 A 20020709 (200254) A61K031-44
EP 1242078 A2 20020925 (200271) DE A61K031-44

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI

KR 2002040767 A 20020530 (200276) A61K031-44

ADT WO 2001008682 A2 WO 2000-EP7356 20000729; CA 2314746 A1 CA 2000-2314746 20000801; AU 2000072717 A AU 2000-72717 20000729; NO 2002000364 A WO 2000-EP7356 20000729, NO 2002-364 20020123; BP 2000012843 A BP 2000 1200

2000-EP7356 20000729, NO 2002-364 20020123; BR 2000012942 A BR 2000-12942 20000729, WO 2000-EP7356 20000729; EP 1242078 A2 EP 2000-960383 20000729, WO 2000-EP7356 20000729; KR 2002040767 A KR 2002-701445 20020201

FDT AU 2000072717 A Based on WO 200108682; BR 2000012942 A Based on WO 200108682; EP 1242078 A2 Based on WO 200108682

PRAI US 1999-147033P 19990803

IC ICM A61K031-44

ICS A61K009-20; A61K045-06; A61P019-00; A61P029-00

AB WO 200108682 A UPAB: 20010402

NOVELTY - The use of flupirtine (2-amino-3-ethoxycarbonylamino-6-(pfluorobenzylamino)pyridine) (I) or its salt is claimed in the treatment of pain (and prevention of development of chronic pain) due to degenerative joint diseases associated with inflammation in cats and dogs.

ACTIVITY - Analgesic; antiarthritic; antirheumatic; antiinflammatory. In analgesic tests in dogs (I) had oral ID50 of 3.5 mg/kg, compared with

18 mg/kg for ibuprofen.

MECHANISM OF ACTION - Activator of noradrenergic descending pathway in the spinal cord; potentiator of antinociceptive GABA-ergic mechanisms; ATP-sensitive potassium ion channel opener; tension-dependent potassium ion channel opener.

USE - (I) is specifically used for treating hip joint dysplasia or pain due to patella dislocation, Dachshund paralysis or cauda-equina syndrome in dogs or cats (all claimed).

ADVANTAGE - (I) has strong analgesic activity and low toxicity and side-effect potential. In particular (I) causes no gastrointestinal, renal or hepatic damage on acute or long-term use (e.g. over 6-12 months). Dwg.0/0

FS CPI

FΑ AB; DCN

MC CPI: B07-D04C; B14-C01; C07-D04C; C14-C01

ABEX

ADMINISTRATION - (I) is formulated in granules, tablets (specifically film, chewable, 2-layer or retard tablets, especially tablets having single or double breakage indentations), bolus, powder, suppositories or injection solution, together with conventional carriers and auxiliaries (especially taste improvers in the case of oral preparations) (all claimed). Unit dose is typically 0.1-20 (preferably 1-5) mg/kg of (I) maleate (to a maximum of 600 mg/day) for oral administration, 0.1-30(preferably 2.5-7.5) mg/kg of (I) maleate (to a maximum of 900 mg/day) for rectal administration or 1.5-5 mg/kg of (I) gluconate by (preferably intramuscular) injection. (I) is optionally used in combination with (a) antiinflammatories, especially selective COX-2 inhibitors (e.g. celecoxib, rofecoxib, valdecoxib or parecoxib), (b) other centrally acting analgesics (e.g. nefopam, tramadol, nalbuphine or dextropropoxyphene), (c) metamizol, (d) antirheumatic agents (e.g. chloroquine, hydroxychloroquine, methotrexate, penicillamine, ademetionine, sulfasalazin or beta-sitosterol), (e) vitamin B (e.g. thiamine, cyanocobalamine) or pyridoxine), (f) steroids (e.g. prednisolone). (g) chondro-protective agents (e.g. chondroitin, glucosamine or polysulfated glycosaminoglycan), (h) TNFalpha-receptors or (i) plant extracts (e.g. devil's claw root, stinging nettle leaf, guaiac wood, willow bark or arnica extract) (all claimed). EXAMPLE - A mixture of 10 kg flupirtine (I), 2.5 kg calcium hydrogen phosphate and 2.5 kg maize starch was granulated with a solution of 1 kg polyvinyl pyrrolidone in 4 kg demineralized water. The granules were blended with 1.3 kg maize starch, 2 kg microcrystalline cellulose, 0.6 kg magnesium stearate, 0.1 kg highly dispersed silica and 1.5 kg Trigarol Digest P (RTM; taste improver), then pressed into 200 mg tablets of diameter of 9 mm and radius of curvature 10 mm, having double breakage indentations. Each tablet contained 100 mg of (I). Breaking strength was $80-100\ N$ and disintegration time was 5 minutes.

L128 ANSWER 7 OF 12 WPIX (C) 2003 THOMSON DERWENT

1999-620282 [53] WPTX

DNC C1999-181043

TΙ Pharmaceutical preparations containing hydrosoluble ketoprofen salts.

DC B03 B05

ΤN GIORGETTI, P L M

PΑ (ERRE-N) ERREKAPPA EUROTERAPICI SPA; (ERRE-N) ERREKAPPA EUROTERAPICI SAS CYC

PΙ WO 9952528 Al 19991021 (199953) * EN 34p A61K031-40

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RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE
          W: CA CN JP KR US
      EP 1024808
                    A1 20000809 (200039) EN
                                                       A61K031-40
          R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE SI
      CN 1263464
                    A 20000816 (200055)
                                                      A61K031-40
      KR 2001013712 A
                       20010226 (200154)
                                                       A61K031-40
      US 6291527
                   B1 20010918 (200157)
                                                       A61K031-19
      JP 2002510336 W 20020402 (200225)
                                                      C07D207-16
                    A5 20021231 (200305)
      CH 692939
                                                      A61K031-19
 ADT WO 9952528 A1 WO 1999-IB626 19990409; EP 1024808 A1 EP 1999-910606
      19990409, WO 1999-IB626 19990409; CN 1263464 A CN 1999-800474 19990409; KR
      2001013712 A KR 1999-711727 19991211; US 6291527 B1 WO 1999-IB626
      19990409, US 1999-445672 19991210; JP 2002510336 W JP 1999-551422
      19990409, WO 1999-IB626 19990409; CH 692939 A5 CH 1998-843 19980411
 FDT EP 1024808 A1 Based on WO 9952528; US 6291527 B1 Based on WO 9952528; JP
      2002510336 W Based on WO 9952528
 PRAI CH 1999-618
                       19990331; CH 1998-843
                                                  19980411
      ICM A61K031-19; A61K031-40; C07D207-16
          A61K009-02; A61K009-06; A61K009-08; A61K009-10; A61K009-12;
           A61K009-16; A61K009-20; A61K009-48; A61K031-192; A61K031-70;
           A61K031-7008; A61P029-00; C07C051-41
 ICA C07C059-84; C07H005-06
 ICI
     C07C059:84
 AB
           9952528 A UPAB: 19991215
     WO
     NOVELTY - Pharmaceutical preparations containing hydrosoluble
     ketoprofen salts obtained by reaction of ketoprofen and
     glucosamine and/or proline and/or hydroxyproline are new.
          DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for
     hydrosoluble salts contained in the pharmaceutical preparations
     characterized by the fact that they are obtained from ketoprofen
     and amino acids, in 0.8-1.2 times the equimolar quantities.
          ACTIVITY - Antiinflammatory; Analgesic. The results obtained for the
     carrageenan edema test in the rat indicate that ketoprofen
     {\tt glucosamine} salt, administered orally at doses of 0.5, 1 and 2
     mg/kg, possesses antiinflammatory activity. The anti-edematogenic effect
     of the test compound is dose-dependent. Ketoprofen
     glucosamine significantly inhibits the reaction process by 30%,
     48% and 72% at oral doses of 0.5, 1 and 2 mg/kg, respectively.
          USE - The preparations are useful for antiinflammatory and analgesic
     treatment of joints and mucous membranes.
          ADVANTAGE - The use of ketoprofen salts is advantageous
     with regard to bioavailability, tolerability and compliance.
     Dwg.0/0
FS
     CPI
FΑ
     AB; DCN
     CPI: B05-A01B; B05-C07; B07-D03; B10-A07; B10-C04B;
MC
          B10-E04B; B14-C01; B14-C03
TECH
                    UPTX: 19991215
     TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Method: The hydrosoluble
     salts are obtained in water solution form and characterized \bar{b}y the fact
     that the synthesis is carried out at neutral pH at 5-60degreesC and that
     the concentration of salts obtained is at least 300 g l to part of -1.
     Alternatively the hydrosoluble salts are in solid form and the synthesis
     is carried out in at least one suitable organic solvent which, after
     reaction, is eliminated at a high temperature and/or reduced pressure.
ABEX
     ADMINISTRATION - Administration is oral (e.g. as tablets, capsules, or
     granules), transdermal, intramuscular, topical or by injection. The
    preparation may be in the form of a solution, irrigation solution,
    mouthwash, suppository, vaginal bougies, gel, cream or foam. No actual
     dosage is given.
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EXAMPLE - An injectable preparation for intramuscular administration

comprised (quantity for 1 unit): ketoprofen glucosamine (170 mg) equivalent to ketoprofen acid (100 mg), benzyl alcohol (90 mg), sodium chloride (27 mg) and water for injectable preparations (up to 3 ml). L128 ANSWER 8 OF 12 WPIX (C) 2003 THOMSON DERWENT **1995-123244** [16] WPIX DNC C1995-056213 Compsn. for improved relief of cold, cold-like and/or flu symptoms contains aminoacid salt of propionic acid non-steroidal antiinflammatory agent with decongestant, expectorant, antihistamine and antitussive. DC B05 MITRA, S IN PΑ (PROC) PROCTER & GAMBLE CO CYC 24 WO 9507103 PΤ A1 19950316 (199516) * EN 18p A61K045-06 RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE W: AU BR CA CN JP PL RU AU 9476040 A 19950327 (199528) A61K045-06 EP 719156 Al 19960703 (199631) EN A61K045-06 R: AT BE CH DE DK ES FR GB GR IE IT LI LU NL PT SE A 19961112 (199651) BR 9407414 W 19970304 (199719) JP 09502201 20p A61K031-045 CN 1130354 A 19960904 (199751) A61K045-06 WO 9507103 A1 WO 1994-US9581 19940824; AU 9476040 A AU 1994-76040 19940824; EP 719156 A1 EP 1994-926020 19940824, WO 1994-US9581 19940824; BR 9407414 A BR 1994-7414 19940824, WO 1994-US9581 19940824; JP 09502201 W WO 1994-US9581 19940824, JP 1995-508695 19940824; CN 1130354 A CN 1994-193312 19940824 AU 9476040 A Based on WO 9507103; EP 719156 A1 Based on WO 9507103; BR 9407414 A Based on WO 9507103; JP 09502201 W Based on WO 9507103 PRAI US 1993-116927 19930907 REP WO 9217171; WO 9217177 TC ICM A61K031-045; A61K045-06 A61K031-085; A61K031-135; A61K031-19; A61K031-195; A61K031-205; A61K031-38; A61K031-40; A61K031-405; A61K031-415; A61K031-42; A61K031-44; A61K031-445; A61K031-485; A61K031-495; A61K031-50; A61K031-505; A61K031-52; A61K031-54; A61K031-55 AB 9507103 A UPAB: 19971222 A compsn. for alleviating cold, cold-like and/or flu symptoms comprises an aminoacid salt of a propionic acid non-steroidal anti-inflammatory agent together with at least one decongestant, one expectorant, one antihistamine and one antitussive. The propionic acid deriv. is pref. ibuprofen, naproxen, benoxaprofen, flurbiprofen, ketoprofen, fenoprofen, fenbufen, indoprofen, pirprofen, carprofen, oxaprozin, pranoprofen, miroprofen, tioxaprofen, suprofen, alminoprofen, or tiaprofen (esp. ibuprofen , naproxen, flurbiprofen, or ketoprofen). The aminoacid salt is pref. triethylamine, tripropylamine, 2-dimethylaminoethanol, 2-diethylaminoethanol, lysine, ornithine, arginine, histidine, caffeine, procain, N-ethylpiperidine, hydrabamine, choline, betaine, ethylenediamine, glucosamine, methylglucamine, theobromine, purine, piperazine or piperidine.

The decongestant is pref. pseudoephedrine, phenylpropanolamine, phenylephrine or ephedrine.

The antitussive is pref. dextromethorphan, chlophedianol,

carbetapentane, caramiphen, noscapine, diphenhydramine, codeine, hydrocodone, hydromorphone or fominoben.

The expectorant is pref. glyceryl guaiacolate, terpin hydrate, ammonium chloride, N-acetylcysteine, bromhexine or ambroxol.

 $\ensuremath{\mathsf{USE}}$ - The compsn. is useful for the treatment of cough, cold-like and/or flu symptoms.

Dosage is 5 to 50 mg of S(+) ketoprofen lysinate, 50 to 800

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mg \ of \ S(+)-ibuprofen lysinate, or 50 to 800 mg \ of \ S(+)-naproxen
      lysinate.
      Dwg.0/0
 FS
      CPI
 FA
      AB; DCN
      CPI: B04-A04; B04-A06; B06-D09; B07-H; B10-A07; B10-A22;
 MC
           B10-B01; B10-B02B; B10-B03B; B10-B04B; B10-C04B;
           B10-C04C; B10-E04B; B14-C03; B14-K01B; B14-K01E
 L128 ANSWER 9 OF 12 WPIX (C) 2003 THOMSON DERWENT
      1992-168661 [21]
                         WPIX
      C1992-077552
 DNC
 ΤI
      New complexes of phenyl-alkanoic acid(s) - esp. ibuprofen of
      naproxen, with amino sugar(s).
 DC
 ΙN
      PARADIES, H H; HASKO, H
      (MEDI-N) MEDICE CHEM-PHARM PUTTER GMBH; (MEDI-N) MEDICE CHEM PHARM FAB
 PA
      PUETTER
 CYC
      24
 PΙ
      EP 486046
                    A2 19920520 (199221)* DE
                                                12p
                                                       C07C057-30
          R: AT BE CH DE DK ES FR GB GR IT LI LU NL SE
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                    Α
                                                10p
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                      19920521 (199229)
                    Α
                                                       C07H005-06
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      CA 2055681
                    Α
                      19920516 (199231)
                                                      C07H005-06
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      HU 59692
                    \mathbf{T}
                       19920629 (199231)
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      BR 9104997
                   A 19920623 (199232)
                                                      C07H005-06
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                   A 19920527 (199306)
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                   A3 19921209 (199344)
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                   A 19931026 (199345)
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     AU 642309
                   B 19931014 (199348)
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     JP 06184003
                   A 19940705 (199431)
                                                9p
                                                      A61K047-48
                   B1 19960501 (199622)
     EP 486046
                                         DE
                                               18p
                                                      C07C057-30
         R: AT BE CH DE DK FR GB IT LI LU NL SE
     DE 59107759
                   G 19960605 (199628)
                                                      C07C057-30
     JP 2542765
                   B2 19961009 (199645)
                                                      A61K047-48
                                                q8
     US 5604206
                   A 19970218 (199713)
                                               g8
                                                      A61K031-70
     EP 486046 A2 EP 1991-119523 19911115; DE 4137683 A DE 1991-4137683
ADT
     19911115; AU 9187904 A AU 1991-87904 19911115; CA 2055681 A CA
     1991-2055681 19911115; HU 59692 T HU 1991-3572 19911114; BR 9104997 A BR
     1991-4997 19911118; ZA 9109075 A ZA 1991-9075 19911115; CS 9103465 A2 CS
     1991-3465 19911115; CN 1061415 A CN 1991-110740 19911115; EP 486046 A3 EP
     1991-119523 19911115; NZ 240616 A NZ 1991-240616 19911115; AU 642309 B AU
     1991-87904 19911115; JP 06184003 A JP 1991-354100 19911115; EP 486046 B1
     EP 1991-119523 19911115; DE 59107759 G DE 1991-507759 19911115, EP
     1991-119523 19911115; JP 2542765 B2 JP 1991-354100 19911115; US 5604206 A
     Cont of US 1991-792479 19911115, US 1994-328722 19940218
    AU 642309 B Previous Publ. AU 9187904; DE 59107759 G Based on EP 486046;
     JP 2542765 B2 Previous Publ. JP 06184003
PRAI DE 1990-4036460 19901115
    No-SR.Pub; 2.Jnl.Ref; DE 2103387; DE 3205077; DE 3639038; DE 3700172; EP
     398288; 1.Jnl.Ref
TC
     ICM A61K031-70; A61K047-48; C07C057-30; C07H000-00
     ICS A61K031-13; A61K031-205; C07C215-10; C07H005-04
ICA A61K031-19; C07H005-06
AB
           486046 A UPAB: 19931213
    Hydrogen-bonded 1:1 complexes (I) of S(+)-phenylalkanoic acids (II) with
    amino sugars (III) are new. (I) are formed by a proton switch interaction
    between the COOH gp. in (II) and the 3-OOH gp. in (III): where R1-R3 are
    not defined. (II) must have a pKa of 3.5-3.9 w.r.t. the COOH gp., and
    (III) must have a pKa of 1.9-4.0 w.r.t. the 3-OH gp..
         USE/ADVANTAGE - (I) are useful as prodrugs of (II), esp. S(+)-
```

ibuprofen (IIa) or S(+)-naproxen, which have analgesic, antiinflammatory, antipyretic and antimicrobial activity. On oral admin., (I) give higher blood levels of (II) in shorter times than (II) alone. 0/2 FS CPI FA AB; GI; DCN CPI: B10-A07; B10-C03; B10-C04B; B10-C04C; MC B12-A01; B12-D01; B12-D07; B12-D08 ABEQ EP 486046 A UPAB: 19931213 $\label{thm:complexes} \mbox{Hydrogen-bonded 1:1 complexes (I) of S(+)-phenylalkanoic acids (II) with } \\$ amino sugars (III) are new. (I) are formed by a proton switch interaction between the COOH gp. in (II) and the 3-OOH gp. in (III): where R1-R3 are not defined. (II) must have a pKa of 3.5-3.9 w.r.t. the COOH gp., and (III) must have a pKa of 1.9-4.0 w.r.t. the 3-OH gp.. USE/ADVANTAGE - (I) are useful as prodrugs of (II), esp. S(+)ibuprofen (IIa) or S(+)-naproxen, which have analgesic, antiinflammatory, antipyretic and antimicrobial activity. On oral admin., (I) give higher blood levels of (II) in shorter times than (II) alone. ABEQ EP 486046 B UPAB: 19960604 Hydrogen-bridge-bound complexes having a stoichiometry of 1:1 comprising S(+)-phenyl alkane acids and amino sugars in which the complex bond is based on interactions of the carboxyl group of the S(+)-phenyl alkane acids and the hydroxyl group at the carbon atom (C3) of the amino sugars having a proton switch of the form (I) and (II); where R1-COOH denotes the S(+)-phenyl alkane acids and (III); denotes the amino sugars, the pKa values relating to the carboxyl group of the S(+)-phenyl alkane acids lying in the range of 3/5-3.9 and the pKa values relating to the hydroxyl group at the carbon atom (C3) of the amino sugars lying in the range of 1.9-4.0. Dwg.1/2 ABEQ US 5604206 A UPAB: 19970326 Preparing a complex of an S(+)-phenyl alkanoic acid and amino sugar comprises: (a) combining the S(+)-phenyl alkanoic acid with an aq. buffer soln. having a pH 5.5-7.5 at 20 deg. C; (b) heating the combined acid and buffer soln. of step (a) to 40 deg. C with constant stirring until a clear transparent soln. is obtd. and all of the S(+)-phenyl alkanoic acid is dissolved; (c) adjusting the pH of the soln. resulting from step (b) to 5.5-6.0 by the addn. of diluted phosphoric acid, then adding the amino sugar in an equimolar amt. relative to the S(+)-phenyl alkanoic acid to form a reaction mixt.; and (d) after complex formation is complete, cooling the reaction mixt. to ppte. therefrom the complex in crystalline form, and recovering the pptd. complex from the reaction mixt. Dwg.1/2 L128 ANSWER 10 OF 12 WPIX (C) 2003 THOMSON DERWENT ΑN 1979-44860B [24] WPIX TΙ Analgesic and antiinflammatory glucosamide deriv. - specifically 2(p-isobutylphenyl)-propionic acid-D-glucosamide. DC PA (HOKR) HOKURIKU PHARM CO LTD CYC JP 54055545 PΙ A 19790502 (197924)* PRAI JP 1977-121533 19771012 A61K031-70; C07H015-18 IC AB JP 54055545 A UPAB: 19930901 A D-glucosamide deriv. of formula (I) is new. It shows analgesic and

anti-inflammatory activity and is useful as a pharmaceutical.

(I) is prepd. by reacting 2-(p-isobutyl-phenyl)propionyl chloride with D-glucosamine. The reaction may be performed by combining a soln. of 2-(p-isobutylphenyl)propionyl chloride in an organic solvent

with an aq. soln. of D-glucosamine which is prepd. by neutralising D-glucosamine hydrochloride with a strong base. Since D-glucosamine free base is unstable, the aq. soln. should be held fairly cooled. The strong base for the neutralisation of Dglucosamine hydrochloride may be NaOH or KOH. Examples of the organic solvent to be used in the reaction include dioxane, chloroform and ether. CPI AΒ CPI: B10-A07; B12-D01; B12-D07 L128 ANSWER 11 OF 12 WPIX (C) 2003 THOMSON DERWENT **1972-55155T** [35] WPIX Compns of glucosamine hydrochloride and anti-rheumatic - agent for treatment of arthrosis and arthritis, with reduced toxicit. (OPFE) OPFERMANN AND SOHN JGW; (OPFE-N) OPFERMANN J G W & CYC 1 DE 2103387 Α (197235)*B 19810730 (198132) DE 2103387 PRAI DE 1971-2103387 19710126 A61K027-14; A61K031-70; A61K045-06 2103387 A UPAB: 19930831 Prepns. contain active components in a molar ratio of 1:10 to 10:1. The anti-rheumatic agents used are mono-, di- or oxyphenylbutazone, indometacin, pyrazolone and/or salicyclic acid. Pref. prepns. contain 0.1-0.5 g. glucosamine HCl, 0.05-0.25 g. anti-rheumatic agent and opt. sulphates and/or iodides of alkali- and/or alkaline earth metals as well as sulphates and/or iodohydrates of organic basis with anti-rheumatic activity. CPI AΒ CPI: B06-D01; B07-D08; B10-A07; B10-C03; B12-D01; B12-D03; B12-D07; B12-D08; B12-D09 L128 ANSWER 12 OF 12 WPIX (C) 2003 THOMSON DERWENT 1966-26657F [00] WPIX Potentiation of neurotropic agents by glucosamine. (TECX) TECPAN SA CYC 1 FR 48047 (196800) *PRAI FR 1965-27377 19650805 48047 M UPAB: 19930831 - Use of glucosamine and salts thereof for potentiation of analgesics, sedatives, muscle relaxants, hypnotics, neuroplegic Preferred potentiator is glucosamine hydrochloride. Typical compounds potentiated are aspirin, paracetamol, phenylbutazone, anti-pyrine, amidopyrine, morphine and its derivatives, chloropromazine, promethazine, barbiturates. Aspirin was tested against phenylbenzoquinone in mice (see Proc. Soc. Exp. Biol. and Med 1965, 118 763; abid. 1957, 95 729) and the ED50 was 130 mg/kg. compared with 19 mg/kg when potentiated with 20% w/w of glucosamine hydrochloride. (a) Tablets contain aspirin 250 mg. glucurono-lactone 100 mg; glucosamine HCl 150 mg. and excipient q.s. Dose 4-6 tabs.

(b) Tablets contain phenobarbitone 10 mg. glucosamine HCl

100 mg. Daily dose 1-4 tablets. FS CPI FA AB

FS

FΑ

MC

TI

DC PΑ

AΒ

FS

FA

MC

ΤI

DC PΑ

MC CPI: **B10-A07**; B12-C07; B12-C08; B12-C09; B12-C10;

daily for treatment of rheumatism

B12-D01; B12-D03; B12-D09; B12-E02

36 S L38, L40

E ANTIARTHRITIC/CT

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=> d his
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(FILE 'HOME' ENTERED AT 10:34:45 ON 27 JAN 2003)
          SET COST OFF
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FILE 'REGISTRY' ENTERED AT 10:34:56 ON 27 JAN 2003
 L1
               3 S 3416-24-8 OR 29031-19-4 OR 66-84-2
                 E .ALPHA.-GLUCOSAMINE/CN
               1 S E4
 L2
 L3
               1 S 6490-70-6
                 E .BETA.-GLUCOSAMINE/CN
               1 S E4
L4
L5
               1 S 14257-69-3
                 E N-ACETYL-D-GLUCOSAMINE/CN
L6
               1 S E3
L7
             297 S (7512-17-6 OR 3416-24-8 OR 6490-70-6 OR 14257-69-3)/CRN
L8
              37 S L7 AND (7664-93-9/CRN OR CLH)
L9
              8 S L8 AND 2/NC
L10
              12 S L1-L6, L9
L11
              1 S IBUPROFEN/CN
L12
              1 S KETOPROFEN/CN
L13
              18 S C13H18O2/MF AND 46.150.18/RID AND 1/NR AND BENZENEACETIC AND
L14
              13 S L13 AND 2 METHYLPROPYL
L15
               3 S L14 AND IBUPROFEN
              15 S L13 NOT L15
L16
L17
              12 S C16H14O3/MF AND 46.150.18/RID AND 2/NR AND BENZENEACETIC AND
L18
               3 S L17 AND KETOPROFEN
               9 S L17 NOT L18
L19
L20
               6 S L11, L12, L15, L18
                 SEL RN
             426 S E1-E6/CRN
L21
L22
               2 S L21 AND L7
     FILE 'HCAPLUS' ENTERED AT 10:44:09 ON 27 JAN 2003
L23
           9874 S L10
          27519 S ?GLUCOSAMINE? OR ACETYLGLUCOSAMINE OR ACETYL (1W) GLUCOSAMINE
L24
L25
          29352 S L23, L24
           8313 S L20
L26
L27
           8906 S IBUPROFEN OR KETOPROFEN
L28
          10023 S L26, L27
L29
           4491 S NSAID
L30
          11691 S (NONSTEROID? OR NON STEROID?) (L) ?INFLAM?
          49473 S ANALGES?
L31
                E ANALGESIC/CT
                E E6+ALL
L32
          27328 S E5
L33
          54026 S E5+NT
                E E22+ALL
L34
          17760 S E5+NT
                E ANTIINFLAM/CT
                E E5+ALL
L35
          19798 S E2
                E E2+ALL
L36
          24177 S E4, E5
L37
          28056 S E3+NT
L38
             36 S L25 AND L28
L39
            329 S L25 AND L29-L37
L40
             23 S L38 AND L39
L41
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E E4+ALL
L42
            4488 S E5, E4+NT
                 E ANTIHISTAMIN/CT
L43
            1153 S E5-E7
                 E E4+ALL
L44
            6793 S E5, E4+NT
                 E MUSCLE RELAXANT/CT
L45
            5669 S E4-E10
                 E E4+ALL
L46
            8857 S E5, E6, E4+NT
                 E DECONGESTANT/CT
L47
             431 S E4, E5
                 E E4+ALL
L48
             431 S E4
                 E BRONCHODIAL/CT
L49
            5474 S E7-E9
                 E E7+ALL
L50
            9708 S E5, E4+NT
L51
           66379 S ANTIARTHRIT? OR ANTIHISTAMIN? OR ANTI() (ARTHRIT? OR HISTAMIN?
L52
             147 S L25 AND L42-L51
L53
             120 S L52 AND L39
L54
              13 S L52 AND L41
L55
              13 S L40 AND L54
                 SEL DN AN 3 10 11
L56
               3 S E1-E9
L57
             131 S L41, L52, L53 AND L23
L58
              26 S L57 AND L26
L59
              23 S L58 NOT L56
L60
              15 S L59 NOT L55
                 SEL DN AN 11
L61
               1 S L60 AND E10-E12
L62
               2 S L22
L63
               6 S L56, L61, L62
                 E RAFFA R/AU
L64
             177 S E4-E9
                 E COWAN A/AU
             236 S E3-E15,E17,E20,E21
L65
                 E TALLARIDA R/AU
L66
             103 S E4-E6
L67
               1 S L64-L66 AND L25
L68
               6 S L63, L67 AND L23-L67
                 SEL HIT RN
     FILE 'REGISTRY' ENTERED AT 11:00:56 ON 27 JAN 2003
L69
             11 S E1-E11
L70
             10 S L69 NOT C16H25NO2
L71
              1 S L69 NOT L70
     FILE 'REGISTRY' ENTERED AT 11:01:43 ON 27 JAN 2003
     FILE 'HCAPLUS' ENTERED AT 11:02:03 ON 27 JAN 2003
     FILE 'REGISTRY' ENTERED AT 11:03:00 ON 27 JAN 2003
L72
             10 S L10, L20 NOT L69
     FILE 'EMBASE' ENTERED AT 11:03:31 ON 27 JAN 2003
L73
          12569 S L25
L74
          18148 S L28
L75
          18174 S ?KETOPROFEN? OR ?IBUPROFEN?
L76
             76 S L73 AND L74,L75
             39 S L76 AND GLUCOSAMINE ?/CT
L77
                E GLUCOSAMINE/CT
                E E40+ALL
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L78
              37 S (GLUCOSAMINE?(L)CB)/CT
 L79
               3 S L77 AND L78
                 E ANALGESIC/CT
                 E E6+ALL
 L80
          280444 S E3+NT
 L81
             291 S L80 AND L73
 L82
              18 S L78 AND L81
 L83
              15 S L82 NOT L79
                 SEL DN AN L83 10
 L84
               1 S E1 AND L83
 L85
             291 S L76, L81
 L86
              78 S L85 AND (COADMIN? OR COMEDIC? OR COPRESCRI? OR COTHERAP? OR C
L87
              15 S L86 NOT AB/FA
L88
              63 S L86 NOT L87
L89
              54 S L88 AND PY<=2001
                 SEL DN AN 5 6 13 30 49
L90
               5 S L89 AND E2-E8
L91
               5 S L84,L90 AND L73-L90
      FILE 'EMBASE' ENTERED AT 11:20:59 ON 27 JAN 2003
     FILE 'WPIX' ENTERED AT 11:21:23 ON 27 JAN 2003
L92
            1860 S L24
                 E GLUCOSAMINE/DCN
                 E E3+ALL
L93
             397 S E2 OR 1615/DRN
L94
              47 S E4
L95
               2 S E6
                 E ACETYLGLUCOSAMINE/DCN
                 E E4+ALL
L96
             135 S E2
L97
            773 S C07H005-06/IC, ICM, ICS
L98
            2655 S L92-L97
L99
           1931 S L24/BIX
L100
           2712 S L98, L99
L101
           1902 S (?KETOPROFEN? OR ?IBUPROFEN?)/BIX
                 E KETOPROFEN/DCN
                 E E3+ALL
            674 S E2
L102
L103
               1 S E4
L104
            278 S E8
L105
              8 S E10
                E IBUPROFEN/DCN
                E E3+ALL
L106
           1621 S E2 OR 1987/DRN
L107
              3 S E4
L108
            478 S E8
L109
             14 S E10
L110
              4 S E14
L111
             28 S E20
L112
             22 S L100 AND L101-L111
L113
             15 S L112 AND M782/M0,M1,M2,M3,M4,M5,M6
L114
              1 S L112 AND (B14-S09 OR C14-S09 OR B12-C09 OR C12-C09)/MC
L115
              1 S L112 AND P861/M0, M1, M2, M3, M4, M5, M6
L116
             15 S L113-L115
L117
             76 S (B14-C01 OR C14-C01 OR B12-D01 OR C12-D01)/MC AND L100
L118
             75 S (P411 OR PR12)/MO,M1,M2,M3,M4,M5,M6 AND L100
L119
             83 S L117, L118
L120
              1 S L112-L119 AND (RAFFA R? OR COWAN A? OR TALLARIDA R?)/AU
L121
             37 S L112-L119 AND (B10-A07 OR C10-A07)/MC
L122
             15 S L112-L119 AND (B10-C04B OR C10-C04B OR B10-C04C OR C10-C04C)/
L123
              6 S L121 AND L122
                SEL DN AN 1 3 4 5
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L125	4 S L123 AND E1-E8 4 S L120,L124 5 S L112-L116,L121-L122 NOT L123-L125
L127	SEL DN AN 7 12 16 18 21 42 44 45 8 S E9-E22 2 S L125,L127 AND L92-L127

FILE 'WPIX' ENTERED AT 11:43:53 ON 27 JAN 2003